SEARCH REQUEST FORM

(347), Sci	entific and Technica	I Information Center		
Requester's Full Name: PATEL Art Unit: (24 Phone N Mail Box and Bldg/Room Location	Tumber 30 6 4 7 6	Y Senai Number: CE) G.
If more than one search is submi	itied, please prioritiz	e searches in order of	need.	d + ++
Please provide a detailed statement of the sinclude the elected species or structures, k utility of the invention. Define any terms known. Please attach a copy of the covers	eywords, synonyms, acron hat may have a special me	yms, and registry numbers, a aning. Give examples or rel	nd combine with the concept or	· · ·
Title of Invention: (D) FN	SED PYRID	INE COMPO	1 MD 3	, .
Inventors (please provide full names):	KOHSHL	JENO et	el. ; pt	¥(
Earliest Priority Filing Date:	0/2/1977	1/4		187 -118
For Sequence Searches Only Please includ appropriate serial number.	e all pertinent information	-	1 / _(5-611-410
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STAFF USE ONLY	Type of Search	Vendors and co	st where applicable	•
Scarcher Hanly	NA Sequence (#)	STN	···	•
Searcher Phone #:	AA Sequence (#)	Dialog	<u> </u>	
Searcher Location:	Structure (#)	Questel/Orbit		•
Date Searcher Picked Up:	Bibliographic	Dr.Link		•
Date Completed: X 5	Litigation	Lexis/Nexis Sequence Systems		
Searcher Prep & Review Time:	Fulltext Patent Family	WWW/Internet		i
Online Time:	Other	Other (specify)		

PTO-1590 (1-2000)

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    (FILE 'HOME' ENTERED AT 15:53:05 ON 05 AUG 2001)
    FILE 'HCAPLUS' ENTERED AT 15:53:30 ON 05 AUG 2001
Ll
          2737 S UENO K?/AU
          1667 S SASAKI A?/AU
L2
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L3
          1166 S OKABE T?/AU
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L5
         14232 S TAKAHASHI K?/AU
L6
             0 S YAMAOTO N?/AU
L7
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Ľ8
           660 S MATSUNAGA M?/AU
           392 S KUBOTA A?/AU
L10
                                                        - inventor search
L11
         34556 S L1-10
             1 S L11 AND CONDENSED PYRIDINE
L12
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L13
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L14
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               SELECT RN L15 1
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            38 S E1-38
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L17
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L18
               STR
            50 S L18
L19
          1960 S LIB FOL parent search
L20
               SAVE L20 PAT850P/A
L21
               STR L18
            50 S L21 SSS SAM SUB=L20
           50 S L21 SSS SAM SUB=L20
826 S L21 SSS FUL SUB=L20 8 2 6 cp ds in Subset search
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L23
               SAVE L23 PAT850S1/A
            15 S L16 AND L23 15 cpdo from L23 are in appl. work (L17)
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             1 S L25 NOT L17 / cite of appl. compounds
L25
L26
L27
            44 S L23
            42 S L27 NOT L25
L28
           40 S L28 AND PY<1999
35 S L29 AND PY<1998 35 C ites w/ PY 21998
7 S L28 NOT L30 = 1 cite is a patent with an aurlie privity date
            40 S L28 AND PY<1999
L29
L30
L31
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=> d que 127
L18

STR

Parent

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N 11

R

VAR G1=CH/N
NODE ATTRIBUTES:
NSPEC IS RC AT 11
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 11

STEREO ATTRIBUTES: NONE
L20 1960 SEA FILE=REGISTRY SSS FUL L18
L21 STR Subset

VAR G1=CH/N
REP G2=(0-6) CH2
VAR G3=CY/14/17/20/22
NODE ATTRIBUTES:
NSPEC IS RC AT 11
CONNECT IS E1 RC AT 20
DEFAULT MLEVEL IS ATOM
GGCAT IS LIN AT 18
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RSPEC I NUMBER OF NODES IS 23

L27

STEREO ATTRIBUTES: NONE

826 SEA FILE=REGISTRY SUB=L20 SSS FUL L21 44 SEA FILE=HCAPLUS ABB=ON PLU=ON L23

=> d bib abs hitstr

- ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2001 ACS 2000:277851 HCAPLUS AN DN 132:313677 Analgesics containing 1-(1-phenethylpiperidin-4-yl)indole, ΤI 1-(piperazin-1-yl)-3-phenylisoquinoline, or 4-(piperazin-1-yl)-6phenylthieno(3,2-c)pyridine derivatives IN Ueno, Kohshi; Sasaki, Atsushi; Kitazawa, Noritaka; Kawano, Koki; Okabe, Tadashi; Takahashi, Keiko: Matsunaga, Manabu: Shinoda, Yukie Eisai Co., Ltd., Japan PCT Int. Appl., 29 pp. so CODEN: PIXXD2 ÐΤ Patent Japanese LA FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE WO 2000023075 WO 1999-JP5761 PΥ A1 20000427 19991019
- W: CA, CN, KR, US
 RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
 PT, SE
 JP 2000191533 A2 20000711 JP 1999-296106 19991019
 PRAI JP 1998-296681 A 19981019
 OS MARPAT 132:313677

$$N-(CH_2)_n$$
 R^3 R^2 I

AB Novel analgesics for various diseases such as headache and migraine and pain and ache in assocn. with trauma, phys. compression, etc. are described. These analgesics, which are useful for the prevention, treatment, or improvement of pains in humans, contain as the active ingredient benzene derivs. represented by general formula (I or II) or pharmacol. acceptable salts thereof (wherein R2, R3 = H, halo, lower alkyl, lower alkoxy, cyano, lower hydroxyalkyl, lower hydroxyalkoxy, N-lower alkylamino, lower alkylsulfonylaminoalkyl; R4 = lower acylaminoalkyl, amido-lower alkylsulfonylaminoalkyl; R4 = lower acylaminoalkyl, hydroxy-lower alkyl; the ring A represents a benzene or thiophene ring). I and II s.c. showed analgesic activity equal to or greater than that of morphine hydrochloride in acetic acid-induced

PATEL 09/852,850

writhing assay in mice. They were also tested for the binding activity to serotonin (5HT) receptor as well as muscle relaxant activity. IT 214611-53-7 214613-26-0 214613-27-1 214613-33-9 214613-49-7 214613-83-9 214613-84-0 214613-89-5 214613-90-8 214618-14-1 223540-38-3 223540-56-5 223540-84-9 223540-90-7 223541-70-6 223542-28-7 223542-29-8 223546-94-9 223546-95-0 223547-08-8 223547-11-3 223547-20-4 223547-21-5 223547-40-8 223547-42-0 223551-27-7 223551-30-2 223557-26-4 265667-20-7 265667-21-8 265667-22-9 265667-23-0 265667-24-1 265667-25-2 265667-26-3 265667-27-4 265667-28-5 265667-35-4 RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (analgesics contg. 1-(1-phenethylpiperidin-4-yl)indole, 1-(piperazin-1-yl)-3-phenylisoquinoline, or 4-(piperazin-1-yl)-6phenylthieno[3,2-c]pyridine derivs.) RN 214611-53-7 HCAPLUS Acetamide, $N-\{[1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-2,3-dihydro-hyd$ CN 1H-indol-6-yl]methyl]- (9CI) (CA INDEX NAME)

$$\begin{array}{c|c} & & & \\ & & & \\ \text{AcNH-CH2} & & & \\ \end{array}$$

RN 214613-26-0 HCAPLUS

CN 1H-Indole-6-acetamide, 1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-N-(2-hydroxyethyl)- (9CI) (CA INDEX NAME)

RN 214613-27-1 HCAPLUS

CN 1H-Indole-6-acetamide, 1-{1-{2-(4-fluorophenyl)ethyl}-4-piperidinyl}-N,N-dimethyl- (9CI) (CA INDEX NAME)

RN 214613-33-9 HCAPLUS

CN 1H-Indole-6-acetamide, 1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-(9CI) (CA INDEX NAME)

$$\bigcap_{H_2N-C-CH_2} \bigcap_{N} \bigcap_{N-CH_2-CH_2-CH_2} \bigcap_{F}$$

RN 214613-49-7 HCAPLUS

CN lH-Indole-6-acetamide, 1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-N-methyl- (9CI) (CA INDEX NAME)

RN 214613-83-9 HCAPLUS

CN lH-Indole-6-acetamide, N-ethyl-1-(1-[2-(2-fluorophenyl)ethyl]-4piperidinyl]- (9CI) (CA INDEX NAME)

$$0 \\ | C \\$$

RN 214613-84-0 HCAPLUS

CN lH-Indole-6-acetamide, 1-[1-[2-(2-fluorophenyl)ethyl]-4-piperidinyl]-N-(2hydroxyethyl)- (9CI) (CA INDEX NAME)

RN 214613-89-5 HCAPLUS

CN Acetamide, N-[[1-[1-[2-(4-methoxyphenyl)ethyl]-4-piperidinyl]-1H-indol-6-yl]methyl]- (9CI) (CA INDEX NAME)

RN 214613-90-8 HCAPLUS

CN Acetamide, N-{[1-[2-(2-fluorophenyl)ethyl}-4-piperidinyl]-1H-indol-6yl]methyl]- (9CI) (CA INDEX NAME)

RN 214618-14-1 HCAPLUS

CN lH-Indole-6-acetamide, 1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-N,N-dimethyl-, ethanedioate (1:1) (9CI) (CA INDEX NAME)

CM :

CRN 214613-27-1 CMF C25 H30 F N3 O

CM 2

CRN 144-62-7 CMF C2 H2 O4

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RN 223540-38-3 HCAPLUS
CN Isoquinoline, 1-(4-ethyl-1-piperazinyl)-3-(4-methoxyphenyl)- (9CI) (CA INDEX NAME)

RN 223540-56-5 HCAPLUS
CN Benzenemethanol, .alpha.-ethyl-4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]- (9CI) (CA INDEX NAME)

RN 223540-84-9 HCAPLUS
CN Benzenepropanol, 4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-methoxy(9CI) (CA INDEX NAME)

RN 223540-90-7 HCAPLUS
CN Ethanol, 2-[4-{1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl)phenoxy]- (9CI)
(CA INDEX NAME)

223541-70-6 HCAPLUS

Benzamide, 4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-N-propyl- (9CI) (CA INDEX NAME) CN

223542-28-7 HCAPLUS RN

1-Propanesulfonamide, N-[[2-chloro-4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]phenyl]methyl]- (9CI) (CA INDEX NAME) CN

223542-29-8 HCAPLUS
1-Propanesulfonamide, N-{[2-chloro-4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]phenyl]methyl]-, dihydrochloride (9CI) (CA INDEX NAME)

●2 HC1

RN 223546-94-9 HCAPLUS
CN Benzenemethanol, 4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl]alpha.-methyl- (9CI) (CA INDEX NAME)

RN 223546-95-0 HCAPLUS
CN Benzenemethanol, 4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl].alpha.-methyl-, ethanedioate (1:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 223546-94-9 CMF C21 H25 N3 O S

CM 2

CRN 144-62-7 CMF C2 H2 O4

RN

223547-08-8 HCAPLUS
2-Propanol, 1-[4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl]phenoxy]- (9CI) (CA INDEX NAME) CN

RN 223547-11-3 HCAPLUS

2-Propanol, 1-[4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl]phenoxy]-, dihydrochloride (9CI) (CA INDEX NAME) CN

●2 HCl

223547-20-4 HCAPLUS

2-Propanol, 1-[4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl]phenoxy]-2-methyl- (9CI) (CA INDEX NAME)

RN

223547-21-5 HCAPLUS
2-Propanol, 1-[4-[4-(4-ethyl-1-piperazinyl)thieno[3,2-c]pyridin-6-yl]phenoxy]-2-methyl-, dihydrochloride (9CI) (CA INDEX NAME) CN

●2 HC1

RN 223547-40-8 HCAPLUS
CN 1-Piperazineethanol, 4-[6-[4-(2-hydroxy-2-methylpropoxy)phenyl]thieno[3,2-c]pyridin-4-yl]- (9CI) (CA INDEX NAME)

RN 223547-42-0 HCAPLUS
CN 1-Piperazineethanol, 4-[6-[4-(2-hydroxy-2-methylpropoxy)phenyl]thieno[3,2-c]pyridin-4-yl]-, dihydrochloride (9CI) (CA INDEX NAME)

●2 HCl

RN 223551-27-7 HCAPLUS
CN Benzenepropanol, 4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-fluoro.alpha.,.alpha.-dimethyl- (9CI) (CA INDEX NAME)

RN 223551-30-2 HCAPLUS
CN Benzenepropanol, 3-{1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl}- (9CI) (CA INDEX NAME)

RN 223557-26-4 HCAPLUS
CN Benzonitrile, 5-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-(2-hydroxyethoxy)- (9CI) (CA INDEX NAME)

RN 265667-20-7 HCAPLUS
CN Acetamide, N-[[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-1H-indol-6yl]methyl]- (9CI) (CA INDEX NAME)

RN 265667-21-8 HCAPLUS

CN 3-Pentanone, 1-[[[1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]-1H-indol-6-yl]methyl]amino]- (9CI) (CA INDEX NAME)

$$\bigcap_{\mathsf{Et-C-CH}_2-\mathsf{CH}_2-\mathsf{NH-CH}_2}^{\mathsf{N}-\mathsf{CH}_2-\mathsf{CH}_2-\mathsf{NH-CH}_2}$$

RN 265667-22-9 HCAPLUS

CN 1H-Indole-6-acetamide, 1-[1-[2-(2-fluorophenyl)ethyl]-4-piperidinyl}-N-methyl- (9CI) (CA INDEX NAME)

$$0 \\ \text{MeNH-} C-CH_2$$

$$N - CH_2 - CH_2$$

RN 265667-23-0 HCAPLUS

CN 1H-Indole-6-acetamide, 1-[1-[2-(2-fluorophenyl)ethyl]-4-piperidinyl](9CI) (CA INDEX NAME)

RN 265667-24-1 HCAPLUS

CN Benzamide, 4-{1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-N-propyl-, monohydrochloride (9CI) (CA INDEX NAME)

• HCl

RN 265667-25-2 HCAPLUS

CN Benzenepropanol, 4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-fluoro-.alpha.,.alpha.-dimethyl-, monohydrochloride (9CI) (CA INDEX NAME) PATEL 09/852,850

● HCl

RN 265667-26-3 HCAPLUS
CN Benzenepropanol, 4-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-methoxy-,
monohydrochloride (9CI) (CA INDEX NAME)

• HCl

RN 265667-27-4 HCAPLUS
CN Benzonitrile, 5-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl]-2-(2-hydroxyethoxy)-, monohydrochloride (9CI) (CA INDEX NAME)

PATEL 09/852,850

• HCl

265667-28-5 HCAPLUS RN

Benzenepropanol, 3-[1-(4-ethyl-1-piperazinyl)-3-isoquinolinyl}-, monohydrochloride (9CI) (CA INDEX NAME) CN

HC1

265667-35-4 HCAPLUS

1H-Indole-6-acetamide, N-acetyl-1-[1-[2-(4-fluorophenyl)ethyl]-4-piperidinyl]- (9CI) (CA INDEX NAME) CN

RE.CNT 4

RE

- (1) Eisai Co Ltd; WO 9843956 Al 1998 HCAPLUS (2) Eisai Co Ltd; WO 9918077 Al 1999 HCAPLUS (3) Meiji Seika Kaisha Ltd; US 5631257 A 1997 HCAPLUS (4) Rhone-Poulenc Rorer S A; US 5563144 A 1996 HCAPLUS

PATEL 09/852.850

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=> d bib abs hitstr 2
L31 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2001 ACS
     1999:521437 HCAPLUS
AN
DN
     131:157754
     Preparation of naphthyridine IL-4 antagonists and G-CSF stimulators
TΙ
     Solomon, Daniel M.; Grace, Michael J.; Fine, Jay S.; Bober, Loretta A.;
IN
     Sherlock, Margaret H.
PA
     Schering Corporation, USA
     U.S., 57 pp.
50
     CODEN: USXXAM
DΤ
     Patent
     English
LA
FAN.CNT 1
     PATENT NO.
                       KIND DATE
                                              APPLICATION NO. DATE
                              19990817
                                              US 1997-878860
                                                               19970619
PΙ
     US 5939431
                        Α
PRAI US 1996-22173
                              19960620
os
     MARPAT 131:157754
     Title compds., e.g., R121NHSO22(NH)a(CO)bR8 (R1 = 3-methyl-2-pyridinyl; Z1
     = 1,7-naphthyridine-6,8-diyl)[I; R8 = alkyl(oxy) or benzyl(oxy); Z =
     phenylene; a,b = 0 or 1) were prepd. as IL-4 antagonists (no data) and G-CSF stimulators. Thus, 8-amino-6-(3-methyl-2-pyridinyl)-1,7-
     naphthyridine was amidated by 4-(AcHN)C6H4SO2C1 to give I (R8 = Me, Z =
     1,4-phenylene, a = b = 1). Data for G-CSF stimulating activity of I were
     given.
IT
     200927-49-7P 200927-65-7P 200927-80-6P
     200928-20-7P 200928-22-9P 200928-24-1P
     RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic
     preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
     (prepn. of naphthyridine IL-4 antagonists and G-CSF stimulators) 200927-49-7 HCAPLUS
RN
     Acetamide, N-[[4-(acetylamino)phenyl]sulfonyl]-N-[6-(3-methyl-2-pyridinyl)-
     1,7-naphthyridin-8-yl]- (9CI) (CA INDEX NAME)
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RN 200927-65-7 HCAPLUS

CN Glycine, N-[(4-aminophenyl)sulfonyl]-N-[6-(3-methyl-2-pyridinyl)-1,7naphthyridin-8-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 200927-80-6 HCAPLUS
CN Acetamide, N-[4-[[methyl[6-(3-methyl-2-pyridinyl)-1,7-naphthyridin-8-yl]amino]sulfonyl]phenyl]- (9CI) (CA INDEX NAME)

RN 200928-20-7 HCAPLUS CN 1,7-Naphthyridine, 8-(1,2-dimethylhydrazino)-6-(3-methyl-2-pyridinyl)-(9CI) (CA INDEX NAME)

RN 200928-22-9 HCAPLUS CN 1,7-Naphthyridine, 8-(1-methylhydrazino)-6-(3-methyl-2-pyridinyl)- (9CI) (CA INDEX NAME)

200928-24-1 HCAPLUS Glycine, N-[$\{4-(acetylamino)phenyl\}sulfonyl\}-N-[6-(3-methyl-2-pyridinyl)-1,7-naphthyridin-8-yl}-, ethyl ester (9CI) (CA INDEX NAME)$ RN CN

RE.CNT 12

RE

- RE
 (2) Anon; GB 1545767 1979 HCAPLUS
 (5) Behrens; US 4942163 1990 HCAPLUS
 (6) De Zwart; J Med Chem 1988, V31, P716 HCAPLUS
 (7) De Zwart; J Med Chem 1989, V32, P487 HCAPLUS
 (8) Demetri; Blood 1991, V78, P2791 HCAPLUS
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

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=> d bib abs hitstr 130 1-35
L30 ANSWER 1 OF 35 HCAPLUS COPYRIGHT 2001 ACS
     1998:25142 HCAPLUS
     128:88786
     Preparation of naphthyridines which affect IL-4 and G-CSF
TI
     Solomon, Daniel M.; Grace, Michael J.; Fine, Jay S.; Bober, Loretta A.;
IN
     Sherlock, Margaret H.
     Schering Corp., USA
PA
     PCT Int. Appl., 98 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
                                              APPLICATION NO. DATE
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                       KIND DATE
                              19971224
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                       A2
     WO 9748368
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              NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UZ, VN, YU,
              AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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     CA 2258752
                        AA
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                                              CA 1997-2222
AU 1997-35673 19970618
                                                                 19970618 <--
                              19980107
                        A1
     AU 9735673
                              19990506
                        A2
     EP 912571
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE,
              LT, LV, FI, RO
                                               CN 1997-197310 19970618
                              19990908
     CN 1228090
                        Α
PRAI US 1996-669185
                              19960620
     WO 1997-US9202
                              19970618
     MARPAT 128:88786
OS
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2.

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

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The title compds. [I; E = II, III, etc.; A = CH, S, N, N(0); L, M, X, Z, W, T, U, V = CH, N, N(0); Y = H, Me; Y1 = H, lower alkyl, Ph, etc.; Q = H, lower alkyl, lower alkyl O(0)CCH2, lower alkyl (0)C; a, b, c, g, h, j = 0-1; f = 1-2; n = 1-6; tt = 0-1; R8 = H, OH, halo, etc.] and their pharmaceutically acceptable salts, useful in the treatment of allergy, inflammation, autoimmune diseases, B-cell lymphomas, tumors, and the after effects of bone marrow transplantation, were prepd. Thus, reaction of 8-amino-6-(3-methyl-2-pyridyl)-1,7-naphthyridine with N-acetylsulfanilyl chloride in the presence of Et3N and DMAP in CH2Cl2 afforded the title compd. IV which resulted in a 4-5-fold increase in G-CSF levels, with an EC50 of 15 .mu.M.
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IT 200927-49-7P 200927-65-7P 200927-80-6P 200928-20-7P 200928-22-9P 200928-24-1P RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of naphthyridines which affect IL-4 and G-CSF)

RN 200927-49-7 HCAPLUS

GI

CN Acetamide, N-[(4-(acetylamino)phenyl]sulfonyl]-N-(6-(3-methyl-2-pyridinyl)-1,7-naphthyridin-8-yl]- (9CI) (CA INDEX NAME)

RN 200927-65-7 HCAPLUS
CN Glycine, N-[(4-aminophenyl)sulfonyl]-N-[6-(3-methyl-2-pyridinyl)-1,7naphthyridin-8-yl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 200927-80-6 HCAPLUS
CN Acetamide, N-[4-[[methyl[6-(3-methyl-2-pyridinyl)-1,7-naphthyridin-8-yl]amino]sulfonyl]phenyl]- (9CI) (CA INDEX NAME)

RN 200928-20-7 HCAPLUS
CN 1,7-Naphthyridine, 8-(1,2-dimethylhydrazino)-6-(3-methyl-2-pyridinyl)(9CI) (CA INDEX NAME)

L30 ANSWER 2 OF 35 HCAPLUS COPYRIGHT 2001 ACS AN 1997:498744 HCAPLUS DN 127:190707

Synthesis and antitumor activity of 3-arylisoquinoline derivatives Cho,-Won-Jea: Yoo. Su-Jeong: Park, Myun-Ji: Chung, Byung-Ho: Lee, TI 'AU

er en 1995 en 1986 en En 1986 en 198

Chong-Ock

College of Pharmacy, Chonnam National University, Kwangju, 500-757, S. cs Korea

Arch. Pharmacal Res. (1997), 20(3), 264-268 CODEN: APHRDQ: ISSN: 0253-6269 Pharmaceutical Society of Korea so

PB

DT Journal

LA English GI

> SEARCHED BY SUSAN HANLEY Phone: 305-4053

Page 3

AB In order to study the structure-activity relationship of 7,8-dimethoxy-2-methyl-3-(4,5-methylenedioxy-2-vinylphenyl)isoquinoline-1(2H)-one (I), which has exhibited significant antitumor activity, Chem. modifications of I were performed to yield the corresponding products, e.g., isoquinoline II. Further systematic uses of an efficient procedure for the synthesis of 3-arylisoquinoline derivs. produced the substituted compds. III (X = H, 4-Br, 4-MeO, 4-Cl, 2-, 3-, 4-Me), which were tested for in vitro antitumor activity against five different human cancer cell lines.

IT 194292-31-4P 194292-32-5P 194292-33-6P 194292-34-7P 194292-35-8P 194292-36-9P 194292-37-0P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (prepn. and antitumor activity of arylisoquinoline derivs.)

RN 194292-31-4 HCAPLUS

CN Isoquinoline, 1-(4-methyl-1-piperazinyl)-3-phenyl-, monohydrochloride (9CI) (CA INDEX NAME)

● HCl

RN 194292-32-5 HCAPLUS

Isoquinoline, 3-(4-bromophenyl)-1-(4-methyl-1-piperazinyl)-,
monohydrochloride (9CI) (CA INDEX NAME)

● HCl

RN 194292-33-6 HCAPLUS
CN Isoquinoline, 3-(4-methoxyphenyl)-1-(4-methyl-1-piperazinyl)-,
monohydrochloride (9CI) (CA INDEX NAME)

HC1

RN 194292-34-7 HCAPLUS
CN Isoquinoline, 3-(4-chlorophenyl)-1-(4-methyl-1-piperazinyl)-,
monohydrochloride (9CI) (CA INDEX NAME)

HCl

● HCl

194292-36-9 HCAPLUS
Isoquinoline, 3-(3-methylphenyl)-1-(4-methyl-1-piperazinyl)-,
monohydrochloride (9CI) (CA INDEX NAME) RN CN

HC1

194292-37-0 HCAPLUS
Isoquinoline, 3-(4-methylphenyl)-1-(4-methyl-1-piperazinyl)-,
monohydrochloride (9CI) (CA INDEX NAME) RN CN

HC1

L30 ANSWER 3 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1996:504767 HCAPLUS 125:275604

AN DN

SEARCHED BY SUSAN HANLEY Phone: 305-4053

Page 6

PATEL 09/852,850

```
DMSO-Ac20 promoted nitration of isoquinolines. One-step synthesis of
TI
      1-nitroisoquinolines under mild conditions
      Baik, Woonphil: Yun, Sangmin; Rhee, Jong Uk; Russell, Glen A.
ΑU
     Dep. Chemistry, Myong Ji Univ., Kyung Ki Do, 449-728, S. Korea J. Chem. Soc., Perkin Trans. 1 (1996), (15), 1777-1779
so
      CODEN: JCPRB4; ISSN: 0300-922X
DТ
      Journal
      English
LA
os
      CASREACT 125:275604
```

Nitroisoquinolines I (R = H, 5-NO2, 4-Br, 3-Me, 5-Me) were directly prepd. AB from the corresponding isoquinolines with potassium nitrite and acetic anhydride in DMSO in good yields.

182184-82-3P IT RL: SPN (Synthetic preparation); PREP (Preparation) (nitration of isoquinolines promoted by potassium nitrite/DMSO/acetic anhydride) 182184-82-3 HCAPLUS RN

Isoquinoline, 3-methyl-1-nitro- (9CI) (CA INDEX NAME)

ANSWER 4 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30 1996:451066 HCAPLUS 125:161858 DN

Selective inhibition of cyclic AMP-dependent protein kinase by ΤI isoquinoline derivatives

Lu, Zhe Xiong; Quazi, Nurul Huda; Deady, Leslie W.; Polya, Gideon M. Sch. Biochem., La Trobe Univ., Victoria, 3083, Australia ΑU

CS

Biol. Chem. Hoppe-Seyler (1996), 377(6), 373-384 SO CODEN: BCHSEI; ISSN: 0177-3593

ηт Journal LA

AΒ

English A large series of isoquinoline derivs. was synthesized including derivs. of isoquinoline, isoquinoline[3,4-c]furazan, 1,2-dihydro-1-oxoisoquinoline, 6-oxopyrimido[1,2-b]isoquinoline, benzo[c][1,8]naphthyridine, pyrazino[2,3-c]isoquinoline and benzimidazo[2,1alisoquinoline as well as further structurally related isoquinoline derivs. and pyrido-2,3-furazans. Representatives of all of these classes of isoquinolines are potent and selective inhibitors of the cAMP-dependent protein kinase (PKA) catalytic subunit (cAK) from rat liver. The most effective cAK inhibitors are a series of 1,3-di-substituted and 1,3,4-tri-substituted isoquinolines (IC50 values 30-50 nm) (compds. Al, A2, A3, A4 and A5) and 2-ethylcarboxy-3-amino-5,6-dihydro-6oxobenzo(c)(1,8)naphthyridine (E1)(IC50 0.08.mu.m). Compds. Al-A5 inhibit CAK in a fashion that is competitive with respect to ATP as substrate. The isoquinoline inhibitors Al-A5 are ineffective or very poor inhibitors of wheat embryo Ca2+-dependent protein kinase (CDPK) and rat brain Ca2+-dependent protein kinase C (PKC), chicken gizzard myosin light chain kinase (MLCK) and potato tuber cyclic nucleotide-binding phosphatase (Pase). El is a moderately effective inhibitor of CDPK and PKC (IC50

PATEL 09/852,850

values 20 and 61 .mu.m, resp.). The bisisoquinoline-1(2H)-one compd. B7
inhibits cAK, CDPK, PKC and MLCK (IC50 values 8, 95, 24 and 7 .mu.m,
resp.) as does J1 [2-(p-bromophenyl)pyrrolo[2,3-c]isoquinoline-5(4H)-one]
(IC50 values 2, 50, 44 and 7 .mu.m, resp.). The very potent
isoquinoline-derived cAK inhibitors found here involve substitution of the
N-contg. isoquinoline ring system and these inhibitors show high
specificity for cAK.
180507-73-7
RL: BAC (Biological activity or effector, except adverse); PRP
(Properties); BIOL (Biological study)
 (selective inhibition of cAMP-dependent protein kinase, other kinases,
 and cyclic nucleotide-binding phosphatase by isoquinoline derivs.)
180507-73-7 HCAPLUS
4-Isoquinolinecarbonitrile, 1-[(4-cyano-3-methyl-1H-2-benzopyran-1ylidene)amino]-3-methyl- (9CI) (CA INDEX NAME)

RN

CN

ANSWER 5 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30 1996:291499 HCAPLUS AN 125:57809 DN Electronic effects in isoquinoline systems TI Zielinski, Wojciech; Kudelko, Agnieszka; Mazik, Monika Institute of Organic Chemistry and Technology, Silesian Technical C5 University, Gliwice, 44-101, Pol. Pol. J. Appl. Chem. (1995), 39(1), 33-38 SO CODEN: PJACE2; ISSN: 0867-8928 ĎΤ Journal English Values of pKa for 1-(N,N-dimethylamino)-3-methylisoquinoline and a series of their 6- and 7-substituted derivs., 3-methylisoquinoline and 1-amino-3-methylisoquinoline were detd. in 50% vol./vol. aq.-methanolic soln. by the spectrophotometric method. The detd. values of pKa and values of pKa for 1-phenyl-3-methylisoquinolines and 1,3dimethylisoquinolines taken from literature were correlated with the Hammett .sigma. consts. Good correlations were obtained for 6-substituted derivs. with .sigma.p consts. and for 7-substituted derivs. with .sigma.m consts. The electronic effects occurring in the studied isoquinoline systems made by substituents present in pyridine and benzene ring are discussed basing on the detd. values. 155999-40-9 155999-41-0 155999-42-1 155999-43-2 155999-44-3 155999-45-4 155999-46-5 177978-22-2 RL: PRP (Properties); RCT (Reactant) (electronic effects in isoquinolines) 155999-40-9 HCAPLUS RN 1-Isoquinolinamine, N,N,3-trimethyl- (9CI) (CA INDEX NAME) CN

RN 155999-41-0 HCAPLUS

N 1-Isoquinolinamine, N,N,3,6-tetramethyl- (9CI) (CA INDEX NAME)

RN 155999-42-1 HCAPLUS

CN 1-Isoquinolinamine, N,N,3,7-tetramethyl- (9CI) (CA INDEX NAME)

RN 155999-43-2 HCAPLUS

CN 1-Isoquinolinamine, 6-chloro-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

RN 155999-44-3 HCAPLUS

CN 1-Isoquinolinamine, 7-chloro-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

RN 155999-45-4 HCAPLUS

CN 1-Isoquinolinamine, 6-methoxy-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

RN 155999-46-5 HCAPLUS

1-Isoquinolinamine, 7-methoxy-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

177978-22-2 HCAPLUS

1-Isoquinolinamine, 7-bromo-N,N,3-trimethyl- (9CI) (CA INDEX NAME) CN

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L30 ANSWER 6 OF 35 HCAPLUS COPYRIGHT 2001 ACS
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1995:869485 HCAPLUS AN

DN 123:343738

Perforated transfer printing media and printing process ТT

Kawakami, Sota; Nakajima, Atsushi; Maejima, Katsumi; Komamura, Tawara IN

Konishiroku Photo Ind, Japan Jpn. Kokai Tokkyo Koho, 52 pp. PA

so CODEN: JKXXAF

ÐΤ Patent

Japanese LA

FAN.

FAN.	CNT 1			•	
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	JP 07172059	A2	19950711	JP 1994-235470	19940929 <
PRA	JP 1993-266507		19931025		

MARPAT 123:343738 OS

The title media, comprising a base sheet, a coloring layer of chelate color formable compd. mixed with binders (e.g., polyvinyl butyral), and color-barrier layer (e.g., gelatins mixed with IR absorbers), are forming perforation on the barrier layer by heat and/or pressure and transfer printing on a printing sheet (e.g., PET film coated with a soln. contg. AB polyvinyl butyral, metallic ion-contg. compd., KF-393, X-22-343). 161581-19-7

RL: MOA (Modifier or additive use); TEM (Technical or engineered material use); USES (Uses)

(chelate azo dyes; perforated transfer printing media and printing process)

161581-19-7 HCAPLUS RN

Benzoic acid, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5-nitro-, ethyl ester (9CI) (CA INDEX NAME) CN

ANSWER 7 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1995:370786 HCAPLUS L30

AN

DN 122:201322

Thermal-transfer recording material using chelating dye ΤI

IN Tanaka, Tatsuo; Kato, Katsunori; Komamura, Tawara

Konishiroku Photo Ind, Japan PA

SO Jpn. Kokai Tokkyo Koho, 13 pp.

CODEN: JKXXAF

DT Patent

Japanese LA

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	JP 06312582	A2	19941108	JP 1993-102714	19930428 <

os MARPAT 122:201322

ĢΙ

- The material contains the dye I(R = substituent on benzene ring; n= 0-3;AB RI= OH, amino) or II (RII = substituent on benzene ring; RI2 = substituent on isoquinoline ring; RI3 = H, halo, monovalent substituent; G = chelatable group; p, q = 0-4)in the thermal-transfer layer. The thermal-transfer layer is contacted with a receptor layer, imagewise heated to form a chelating dye by the reaction of the dye with a metal ion to give images. The materials show good storage stability, and give high d. cyan images.
- 161581-12-0 161581-13-1 161581-14-2

161581-19-7

RL: DEV (Device component use); USES (Uses)

(thermal-transfer recording material contg. chelating dye)

161581-12-0 HCAPLUS RN

4-Isoquinolinol, 1-[(5-chloro-2-hydroxy-3-nitrophenyl)azo]-3-methyl- (9CI) CN (CA INDEX NAME)

RN 161581-13-1 HCAPLUS
CN 4-Isoquinolinol, 1-[[2-hydroxy-5-(methylsulfonyl)-3-nitrophenyl]azo]-3-methyl- (9CI) (CA INDEX NAME)

RN 161581-14-2 HCAPLUS
CN Benzonitrile, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5nitro- (9CI) (CA INDEX NAME)

RN 161581-19-7 HCAPLUS
CN Benzoic acid, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5nitro-, ethyl ester (9CI) (CA INDEX NAME)

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ANSWER 8 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1995: 339706 HCAPLUS
L30
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AN

DN 122:174514

Thermal-transfer recording material and recording method by chelation Kato, Katsunori; Tanaka, Tatsuo; Komamura, Tawara TI

IN

Konishiroku Photo Ind, Japan

Jpn. Kokai Tokkyo Koho, 12 pp.

CODEN: JKXXAF

DΤ Patent

Japanese LA

FAN.	CNT 1			•	
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PΙ	JP 06316164	A2	19941115	JP 1993-106738	19930507 <
OC.	MADDAT 122-17/51	А			

GI

$$\begin{array}{c|c}
OH & & & \\
N = N & & \\
OR & & & \\
NC & & \\
NC & & \\
NC & & \\
NC & & & \\
NC$$

- The material contains an azo dye I [R = (substituted) alkyl, cycloalkyl; A = (substituted) 5- or 6-membered ring, 9- or 10-membered condensed ring] or II [R1, R2 = H, substituent; A = (substituted) 6-membered ring, AΒ condensed ring) in a transfer layer on a substrate. Images are formed by thermal chelating reaction of the azo dye with a metal ion. High-d. and stable cyan images are obtained.
- 161195-94-4 161195-95-5

RL: DEV (Device component use); RCT (Reactant); USES (Uses) (thermal-transfer recording material contg. azo chelating dye for cyan image)

161195-94-4 HCAPLUS RN

Propanedinitrile, [[2-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-CN

5-nitrophenyl]methylene]- (9CI) (CA INDEX NAME)

RN 161195-95-5 HCAPLUS

CN Propanedinitrile, [[2-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]5-(trifluoromethyl)phenyl]methylene]- (9CI) (CA INDEX NAME)

L30 ANSWER 9 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1995:237241 HCAPLUS

DN 122:81247

- TI A short facile route to 1-hydrazinoisoquinoline: Ring closure reactions of substituted 1-hydrazinoisoquinoline derivatives and substituted 2-(4-carbethoxy)phenyl-1(2H)-isoquinolinone derivatives and their biological activity
- AU Pinto de Souza, Eleanor; Fernandes, Peter S.

CS NSR Lab., St. Xavier's Coll., Bombay, 400 001, India

SO Indian J. Chem., Sect. B: Org. Chem. Incl. Med. Chem. (1994), 33B(12), 1150-8
CODEN: IJSBDB; ISSN: 0376-4699

DT Journal

LA English

AB A short facile synthesis of 1-hydrazinoisoquinoline from 1-chloroisoquinoline is reported. Substituted 1,2,4-triazolo[3,4-a]isoquinolines were prepd. from 1-hydrazino-7-methoxy-3-methylisoquinoline. The compd. underwent cyclization with acetic anhydride, benzoyl chloride, di-Et malonate, benzoin, nitrous acid, acetylacetone, Et acetoacetate and di-Et acetylenedicarboxylate. Substituted 2-[4-(4-amino-5-mercapto-1,2,4-triazol-3-yl)phenyl]-1(2H)-isoquinolinone was prepd. Furthermore, 2-[4-(s-triazolo[3,4-b)[1,3,4]thiadiazol-3-yl)phenyl]-1(2H)-isoquinolinone and

2-[4-(s-triazolo[3,4-b][1,3,4]thiadiazin-3-yl)phenyl]-1(2H)-isoquinolinonewere prepd. All the compds. have been tested for their antibacterial activity; by the agar method all compds. were inactive at 50 .mu.g per

160518-59-2P 160518-60-5P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of)

RN

160518-59-2 HCAPLUS Isoquinoline, 1-(3,5-dimethyl-1H-pyrazol-1-yl)-7-methoxy-3-methyl- (9CI) CN (CA INDEX NAME)

160518-60-5 HCAPLUS

Pyrano[2,3-c]pyrazol-6(1H)-one, 3a,7a-dihydro-1-(7-methoxy-3-methyl-1-isoquinolinyl)-4-methyl- (9CI) (CA INDEX NAME)

ANSWER 10 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30

1994:457449 HCAPLUS

121:57449

Syntheses of 2,4-diaminopyrimidines and 1-aminoisoquinolines in the reactions of alkyl and benzyl ketones with cyanamide and N, N-dimethylcyanamide

Zielinski, Wojciech; Mazik, Monika ΑŲ

Inst. Org. Chem. Technol., Silesian Tech. Univ., Gliwice, 44-101, Pol. Heterocycles (1994), 38(2), 375-82 CODEN: HTCYAM; ISSN: 0385-5414 CS

SO

DТ Journal

English LA

GΙ

The reaction of alkyl and benzyl ketones with cyanamide and

PATEL 09/852,850

N,N-dimethylcyanamide in the presence of POCl3 was examd. At the first stage, chloroformamidine derivs, were formed. In the presence of TiCl4, they underwent further reactions to give derivs. of 1-aminoisoquinoline I (R2 = Ph, substituted Ph) and 2,4-diaminopyrimidine II (R1 = alkyl, Ph, substituted Ph). The effect of constitution of substrates on adequate ratios of heterocyclic compds. is discussed.

155999-40-9P 155999-41-0P 155999-42-1P 155999-43-2P 155999-44-3P 155999-45-4P 155999-46-5P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 155999-40-9 HCAPLUS RN

1-Isoquinolinamine, N,N,3-trimethyl- (9CI) (CA INDEX NAME) CN

155999-41-0 HCAPLUS RN

1-Isoquinolinamine, N,N,3,6-tetramethyl- (9CI) (CA INDEX NAME)

155999-42-1 HCAPLUS RN

1-Isoquinolinamine, N,N,3,7-tetramethyl- (9CI) (CA INDEX NAME) CN

155999-43-2 HCAPLUS RN

1-Isoquinolinamine, 6-chloro-N,N,3-trimethyl- (9CI) (CA INDEX NAME) CN

155999-44-3 HCAPLUS RN

1-Isoquinolinamine, 7-chloro-N,N,3-trimethyl- (9CI) (CA INDEX NAME) CN

RN 155999-45-4 HCAPLUS

1-Isoquinolinamine, 6-methoxy-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

CN

RN 155999-46-5 HCAPLUS

CN 1-Isoquinolinamine, 7-methoxy-N,N,3-trimethyl- (9CI) (CA INDEX NAME)

L30 ANSWER 11 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1994:244949 HCAPLUS

DN 120:244949

TI New syntheses of heterocycles with vinyl- and divinylcarbodiimides: pyrroles, triazoles, pyrimidines, pyrindines, isoquinolines and thiazolylisothiazoles

AU Capuano, Lilly; Hammerer, Volker; Huch, Volker

CS Fachbereich 11.2, Org. Chem., Univ. Saarlandes, Saarbruecken, D-66041, Germany

SO Liebigs Ann. Chem. (1994), (1), 23-7 CODEN: LACHDL; ISSN: 0170-2041

DT Journal

LA German

OS CASREACT 120:244949

GI

The title compds. I and R2CH:CR1N:C:NCR1:CHR2 [II, R1 = Ph, 4-ClC6H4, 2-naphthyl, 4-MeC6H4; R2 = Ph, 4-MeC6H4] react with diazomethane either by loss or by retention of the diazo nitrogen, to afford 3,4-dihydro-2-imino-2H-pyrroles or vic-triazoles, resp. The [4 + 2] addn. of benzylidenemethylamine or alicyclic enamines to II gives partially hydrogenated pyrimidine, pyrindine or isoquinoline. Thermolysis of II proceeds with spontaneous dehydrogenation, giving high yields of 1-(1-indolyl)isoquinolines. The pyrrole III, when melted with sulfur, undergoes both dehydrogenation and sulfur insertion, whereby the hitherto unknown thiazolylisothiazole IV is formed. Its structure has been elucidated by an x-ray diffraction anal. A synthesis of

2,3,5-triphenylimidazo[2,1-a]isoquinoline is reported. 154421-00-8P 154421-01-9P IT RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 154421-00-8 HCAPLUS

RN Isoquinoline, 3-phenyl-1-(2-phenyl-1H-indol-1-yl)- (9CI) (CA INDEX NAME)

154421-01-9 HCAPLUS ŔŊ

Isoquinoline, 3-(4-methylphenyl)-1-[2-(4-methylphenyl)-1H-indol-1-yl]-CN (9CI) (CA INDEX NAME)

ANSWER 12 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1992:573507 HCAPLUS ΑN

DN 117:173507

Thermal-transfer recording materials and recording therewith ΤI

Miura, Akio; Komamura, Tawara; Nakayama, Noritaka Konica K. K., Japan IN

PA

Jpn. Kokai Tokkyo Koho, 9 pp. CODEN: JKXXAF SO

ÐΤ Patent

LA Japanese

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE JP 1990-203739 PΙ JP 04089288 19920323 19900731 <-os MARPAT 117:173507

The title materials providing lightfast high-d. cyan images by chelation with metal ions in the receptor contain a layer contg. cyan dyes I (A1-2 = electron withdrawing group; G = chelating group; Q = a group of atoms

PATEL 09/852,850

forming 5- or 6- membered heterocyclic ring), e.g., thermally diffusible

108831-03-4 108831-05-6 143587-62-6 IT

RN

RL: USES (Uses)
(dye, cyan, for thermal transfer recording inks)
108831-03-4 HCAPLUS
4-Isoquinolinol, 1-[[2-hydroxy-3-nitro-5-(trifluoromethyl)phenyl]azo]-3-methyl- (9CI) (CA INDEX NAME) CN

108831-05-6 HCAPLUS
4-Isoquinolinol, 1-[[2-hydroxy-3,5-bis(trifluoromethyl)phenyl]azo]-3-methyl- (9CI) (CA INDEX NAME) CN

143587-62-6 HCAPLUS RN Benzonitrile, 5-chloro-2-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]- (9CI) (CA INDEX NAME)

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ANSWER 13 OF 35 HCAPLUS COPYRIGHT 2001 ACS
1.30
     1990:611863 HCAPLUS
AN
DN
     113:211863
     Preparation of 1(2H)-isoquinolones and 1-isoquinolineamines as neoplasm
TI
     inhibitors
     Behrens, Carl H.
     du Pont de Nemours, E. I., and Co., USA
PA
     U.S., 13 pp.
     CODEN: USXXAM
DΤ
     Patent
LA
     English
FAN.CNT 1
                                            APPLICATION NO.
                                                             DATE
     PATENT NO.
                      KIND
                             DATE
                                                             19890307 <--
                             19900717
                                            US 1989-322191
     US 4942163
                       A
     MARPAT 113:211863
os
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$$R^{2}$$
 R^{3}
 R^{4}
 R^{1}
 R^{R}
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 R^{7}
 R^{1}

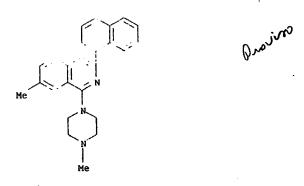
The title compds. [I and II; R = 1-naphthyl; R1, R2, R4 = H, Me, C1; R3 = H, alkyl, C1, NR52, N+R53I-; R5 = H, alkyl; R6, R7 = H, alkyl, (CH2)nNR52; NR6R7 = piperazino, 4-alkylpiperazino; n = 2-8] were prepd. Thus, 5-nitro-N,N,2-trimethylbenzamide (prepn. given) was hydrogenated over Pd/C and the product stirred overnight with 2n-modified NaBH3CN in MeOH contg. HCHO to give 2,4-Me(Me2N)C6H4CONMe2 which was stirred 1 h at -78.degree. with (Me2CH) 2NLi in THF followed by addn. of 1-cyanonaphthalene and stirring for 3 h to give, after acidification, I.HCl (R1 = R3 = R4 = H, R2 = NMe2). The latter increased survival time of mice inoculated with L1210 murine leukemia cells by 156% over controls at 6 mg/kg/day for 9 days. 130370-12-6P 130370-14-8P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of, as neoplasm inhibitor)

130370-12-6 HCAPLUS RN

Isoquinoline, 7-methyl-1-(4-methyl-1-piperazinyl)-3-(1-naphthalenyl)-CN (9CI) (CA INDEX NAME)



130370-14-8 HCAPLUS

1,2-Ethanediamine, N,N,N'-trimethyl-N'-(7-methyl-3-(1-naphthalenyl)-1isoquinolinyl] - (9CI) (CA INDEX NAME)

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CH2-CH2-NMe2
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ANSWER 14 OF 35 HCAPLUS COPYRIGHT 2001 ACS
L30
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1987:431094 HCAPLUS NΑ

107:31094 DN

TI Color photographic recording material

Bergthaller, Peter; Schenk, Guenther; Wolfrum, Gerhard; Runzheimer, Hans Volker: Heidenreich, Holger

PA Agfa-Gevaert A.-G., Fed. Rep. Ger.

SO. Ger. Offen., 81 pp. CODEN: GWXXBX

DΤ Patent

LA German

PAN.CNI I				
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE .
PI DE 3107540	A1	19820916	DE 1981-3107540	19810227 <
EP 59354	A1	19820908	EP '1982-101076	19820213 <
EP 59354	B1	19840620		
R: BE, DE,	FR, GB			,
US 4418143	A	19831129	US 1982-351103	19820222 <
JP 57158637	A2	19820930	JP 1982-31647	19820227 <
JP 03068371	B4	19911028		
PRAI DE 1981-3107540		19810227		•

GΙ

For diagram(s), see printed CA Issue. Diffusable azo dyes of the formula I (R, Rl = electroneg. substituents AB whose meta sigma value .delta.m satisfies .gtoreq.1 of the relations .sigma.m(R), .sigma.m(R1) .gtoreq. +0.33; .sigma.m(R) + .sigma.m(R1) .sigma.m(R), .sigma.m(R) .gtoreq. +0.33; .sigma.m(R) + .sigma.m(R), represented the sigma.m(R) and R1 = SO2R3 where R3 = M, OH, NH2, NHR4 where R4 = alkyl, aryl, alkylsulfonyl, arylsulfonyl, or acyl; R2 = a chelate-forming group; A = 2-amino-3-hydroxypyridine, a 4,5-diphenylimidazole, or a 4-hydroxyisoquinoline ring) are described which are freed upon imagewise development from the corresponding dye releaser and form blue or cyan metal-dye complexes. The dyes, which are useful in color diffusion-transfer photog. materials, give esp. clear cyan color tones when complexed with Ni and Cu complexes. polyethylene-coated paper was coated with a red-sensitized gelatin-Ag(Br,I) emulsion contg. an electron donor compd., a dye releaser of the formula II, and an oil former, a protective layer, and a hardening layer. This element was then exposed through a step wedge, combined with a receptor sheet, and then processed to give a dye image with a Dmin of 0.2, a Dmax of 1.9, a relative sensitivity of 85, and a d. loss of 15% when exposed to a Xe light (4.8 .times. 106 lx-h).

IT 108831-02-3

RL: RCT (Reactant)

(acetylation and chlorination of)

RN 108831-02-3 HCAPLUS

Benzenesulfonamide, 4-hydroxy-3-{(4-hydroxy-3-methyl-1-isoquinolinyl)azo}-5-nitro- (9CI) (CA INDEX NAME)

108830-92-8 RL: USES (Uses)

(photog. azo dye-releasing compd.) 108830-92-8 HCAPLUS

RN

Benzenesulfonamide, N-{5-[[1-{4,5-dimethyl-3,6-dioxo-2-propyl-1,4-cyclohexadien-1-yl)tetradecyl]sulfonyl}-2-methylphenyl}-4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo}-5-nitro-(9CI) (CA INDEX NAME) CN

IT

108831-15-8P . RL: PREP (Preparation)

(prepn. and reaction of diazotized) 108831-15-8 HCAPLUS

2-Naphthalenecarboxamide, 4-[[[3-chloro-4-hydroxy-5-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]phenyl]sulfonyl]amino]-1-hydroxy-N,N-dioctadecyl- (9CI)

IT 108831-13-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and reaction of, with cetyloxyphenylaminoindole) 108831-13-6 HCAPLUS

RN

CN Benzenesulfonyl chloride, 4-(acetyloxy)-3-[[4-(acetyloxy)-3-methyl-1isoquinolinyl]azo]-5-nitro- (9CI) (CA INDEX NAME)

ΙT 108831-00-1P 108831-03-4P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 108831-00-1 HCAPLUS RN

Benzenesulfonamide, 3-chloro-4-hydroxy-5-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]- (9CI) (CA INDEX NAME)

RN 108831-03-4 HCAPLUS
CN 4-Isoquinolinol, 1-[[2-hydroxy-3-nitro-5-(trifluoromethyl)phenyl]azo]-3-methyl- (9CI) (CA INDEX NAME)

108830-95-1 HCAPLUS CN

2-Naphthalenecarboxamide, N-[4-[2,4-bis(1,1-dimethylpropyl)phenoxy]butyl]-1-hydroxy-4-[[[4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5-nitrophenyl]sulfonyl]amino]- (9CI) (CA INDEX NAME)

IT 108830-99-5D, nickel complex 108831-00-1D, copper and nickel complexes 108831-01-2D, nickel complex 108831-02-3D, copper and nickel complexes 108831-03-4D, nickel complex 108831-04-5D, copper and nickel complexes 108831-05-6D, nickel complex 108859-46-7D, nickel complex

RL: PRP (Properties) (spectral properties of, color photog. applications in relation to) 108830-99-5 HCAPLUS

RN

Benzenesulfonamide, 5-chloro-2-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]- (9CI) (CA INDEX NAME) CN

RN 108831-00-1 HCAPLUS

Benzenesulfonamide, 3-chloro-4-hydroxy-5-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]- (9CI) (CA INDEX NAME)

RN 108831-01-2 HCAPLUS

Benzenesulfonic acid, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5-nitro- (9CI) (CA INDEX NAME)

108831-02-3 HCAPLUS RN

Benzenesulfonamide, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5-nitro-(9CI) (CA INDEX NAME) CN

RN 108831-03-4 HCAPLUS CN 4-Isoquinolinol, 1-[[2-hydroxy-3-nitro-5-(trifluoromethyl)phenyl]azo]-3-methyl- (9CI) (CA INDEX NAME)

RN 108831-04-5 HCAPLUS
CN 1,3-Benzenedisulfonamide, 4-hydroxy-5-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]- (9CI) (CA INDEX NAME)

RN 108831-05-6 HCAPLUS
CN 4-Isoquinolinol, 1-[[2-hydroxy-3,5-bis(trifluoromethyl)phenyl]azo]-3-methyl- (9CI) (CA INDEX NAME)

RN 108859-46-7 HCAPLUS

CN Benzenesulfonic acid, 3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-4[(methylsulfonyl)amino]-5-nitro- (9CI) (CA INDEX NAME)

L30 ANSWER 15 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1986:604283 HCAPLUS

DN 105:204283

TI Cytoplasmic vacuolation of pancreatic .beta. cells of rats after oral administration of a derivative of isoquinoline

AU Kast, A.; Ueberberg, H.

CS Dep. Exp. Pathol., Nippon Boehringer Ingelheim Co., Ltd., Yato, Japan

SO Toxicol. Appl. Pharmacol. (1986), 85(2), 274-85 CODEN: TXAPA9; ISSN: 0041-008X

DT Journal

LA English

GI

AB Islet of Langerhans .beta.-cells were studied in Sprague-Dawley rats dosed

PATEL 09/852,850

by gavage with 0 (control), 75, 150, 250 or 300 mg/kg/day SH 966BS (I) 58138-24-2]. All doses caused a significant and dose-dependent increase in serum glucose (diabetes mellitus). At 250 mg/kg, degranulation of .beta.-cells was discovered after 1 day and vacuole formation after 2 days. Ultrastructural alterations compared well with that seen after treatment with cyproheptadine and other structurally related compds. The vacuolation of .beta.-cells was fully developed following 6 wk of daily treatment, when a dose-dependent elevation of blood glucose was 1st obsd. The effects were more severe in males than in females. Lesions were reversible within 6 wk except at 300 mg/kg in males.

. IT · 58138-24-2

> RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (toxicity of, to pancreas .beta.-cells, cytoplasmic vacuolation response to)

58138-24-2 HCAPLUS RN

Isoquinoline, 1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)- (9CI) (CA CN INDEX NAME)

ANSWER 16 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30

1986:88467 HCAPLUS AN

DN 104:88467

ΤI Central nervous system active compounds. XV. 2-Arylisoxazol-5(2H)-ones

AII Hung, Tran V.; Janowski, Wit K.; Prager, Rolf H.

CS Dep. Org. Chem., Univ. Adelaide, Adelaide, 5001, Australia

so Aust. J. Chem. (1985), 38(6), 931-7 CODEN: AJCHAS: ISSN: 0004-9425

DТ Journal

LA English

os CASREACT 104:88467

GI

Et 5-oxo-2,5-dihydroisoxazole-4-carboxylate was treated with a no. of If (R = isoquinolinyl, quinolinyl, purinyl, pyrimidinyl, pyridinyl, pyridazinyl, benzothiazolyl, quinazolinyl, triazinyl). I generally cause loss of motor control in mice, but are relatively toxic.

ΙT 100422-70-6P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (prepn. and central nervous system activity of)

100422-70-6 HCAPLUS RN

4-Isoxazolecarboxylic acid, 2,5-dihydro-2-(3-methyl-1-isoquinolinyl)-5-oxo-CN , ethyl ester (9CI) (CA INDEX NAME)

ANSWER 17 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30

ΔN 1985:561857 HCAPLUS

DN 103:161857

ΤI Photoisomerization and relaxation of symmetrical triazacarbocyanine dyes in an alcohol mixture

Balli, Heinz; Eichenberger, Thomas; Hellrung, Bruno; Scheibli, Peter

Inst. Farbenchem., Univ. Basel, Basel, CH-4056, Switz. Helv. Chim. Acta (1985), 68(5), 1394-400

CODEN: HCACAV; ISSN: 0018-019X

DT Journal

LA German

GI

$$\begin{array}{c|c}
R & X & N = NN \\
R^1 & N^+ & Et & BF_4 & Et \\
\end{array}$$

Photoisomerization of I (R = H, Br, NH2, NO2) and of II (R = H and R1 = H, Me or RR1 = benzo, 1,2-naphtho; X = S, Se, CH:CH, o-C6H4) in 90:5:5 EtOH-MeOH-iso-PrOH at 110-250 K was followed by a 1st-order thermal reverse isomerization in the dark. For II (R = R1 = H, X = CH:CH) [2805-63-2] the irradn. resulted in a decrease in visible absorption intensity with no shift in .lambda.max, whereas most of the other II showed a hypsochromic shift of .lambda.max accompanied by a decrease in intensity. For II (R = R1 = H, X = o-C6H4) [3801-71-6] and 3 other II, irradn. resulted in a shift in the ratio of intensities of 2 absorption bands. With I the electron-donor substituents (OMe, NH2) increased the rate of the dark reaction and NO2 groups decreased the rate. The mechanism proposed involves cis-trans isomerization around the N:N bond, by inversion after partial rotation.

98621-70-6 RL: USES (Uses)

(photoisomerization and subsequent thermal reversion of, kinetics and mechanism of)

98621-70-6 HCAPLUS RN

CN Isoquinolinium, 2-ethyl-1-(3-(2-ethyl-3-methyl-1(2H)-isoquinolinylidene)-1triazenyl)-3-methyl-, tetrafluoroborate(1-) (9CI) (CA INDEX NAME)

CM

CRN 98621-69-3 CMF C24 H26 N5

CM 2

CRN 14874-70-5 CMF B F4 CCI CCS

L30 ANSWER 18 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1985:87554 HCAPLUS

DN 102:87554

TI Silver halide color photographic materials

A Fuji Photo Film Co., Ltd., Japan

O Jpn. Kokai Tokkyo Koho, 23 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

PI JP 59154448 A2 19840903 JP 1983-28927 GI For diagram(s), see printed CA Issue.

AB Ag halide color photog. photosensitive materials contain azo dye forming compd. I or II (A = heterocycle; B = 5-membered heterocycle; Z = bond, divalent moiety; R = group which is sepd. from the azo dye or its precursor during processing by an alk. soln.; Rl, R2 = H, or a substituent which does not exhibit photog. degrdn. effects). The dyes formed from I or II form stable chelates with metals, and hence useful for forming stable images in image receptor layer. Thus, a polyester film support was coated with (1) a dye-mordanting layer contg. Ni acetate and divinylbenzene-N-methyl-N-(vinylbenzyl)piperidinium chloride-styrene copolymer, (2) a layer contg. acylamide-Na N-vinylbenzyliminodiacetate copolymer, (3) a reflector layer contg. TiO2, (4) a carbon black-contg. layer, (5) a layer contg. III, (6) a red-sensitive internal latent image type Ag halide emulsion layer, (7) a gelatin layer contg. 2,5-di-tert-pentadecylhydroquinone, and (8) an overcoat layer to give a diffusion transfer photosensitive film. The diffusion transfer photog. film gave images with azo dye-Ni chelate (having .gamma.max 650) with high Dmax and low Dmin.

IT 94767-38-1P RL: PREP (Preparation) 19830223 <--

(prepn. of, as diffusion-transfer photog. dye-releasing compd.)
RN 94767-38-1 HCAPLUS
CN 1H-Indazole-5-sulfonamide, N-[5-(1,1-dimethylethyl)-4-(hexadecyloxy)-2-

hydroxyphenyl]-7-{(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-4-nitro- (9CI) (CA INDEX NAME)

IT 94737-95-8 RL: RCT (Reactant)

(reaction of, with amino-tert-butylhexadecyloxyphenol hydrochloride)

RN 94737-95-8 HCAPLUS

CN lH-Indazole-5-sulfonyl chloride, 7-[[4-(benzoyloxy)-3-methyl-1-isoquinolinyl]azo]-4-nitro- (9CI) (CA INDEX NAME)

L30 ANSWER 19 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1982:615949 HCAPLUS

DN 97:215949

TI A synthesis of alkylated 3-aminoisoquinolines and related compounds

AU Liepa, Andris J.

CS Div. Appl. Org. Chem., CSIRO, Melbourne, 3001, Australia SO Aust. J. Chem. (1982), 35(7), 1391-403

O Aust. J. Chem. (1982), 35(7), 1391-40 CODEN: AJCHAS; ISSN: 0004-9425

DT Journal

LA English

AB N,N-Dialkyl derivs. of 3-aminoisoquinoline have been prepd. by reaction of nitriles with various arylacetic acid tertiary amides in the presence of POC13. The synthesis has been extended to include a benzoisoquinoline and annulated isoquinolines by the selection of appropriate amide and nitrile

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precursors.
IT
     83814-30-6P
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RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 83814-30-6 HCAPLUS RN

CN 1-Isoquinolinamine, 6,7-dimethoxy-N,N-dimethyl-3-(4-morpholinyl)- (9CI) (CA INDEX NAME)

ANSWER 20 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1981:569009 HCAPLUS

DN 95:169009

Isoquinoline acetic acids and pharmaceutical compositions containing them TI

Schnur, Rodney Caughren Pfizer Inc., USA IN

PA

SO Eur. Pat. Appl., 24 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.	CNT	1				
	PAT	ENT NO.	KIND	DATE	APPLICATION NO.	DATE
PΙ	EΡ	30861	A2	19810624	EP 1980-304517	19801215 <
	EΡ	30861	A3	19810923		
	EP	30861	B1	19830727		
		R: BE, CH,	DE, FR	, GB, IT, LU	, NL, SE	
	US	4283539	A	19810811	US 1979-104939	19791218 <
	JP	56092871	A2	19810727	JP 1980-177066	19801215 <
	JР	62010508	B4	19870306		
	DK	8005364	A	19810619	DK 1980-5364	19801217 <
	DK	149569	В	19860728		
	DK	149569	С	19870202		
PRAI	US	1979-104939		19791218		
GI						

- Acids and esters I, II, and III [R = H, Me; Rl = H, alkyl; R2 = (un) substituted benzyl or benzyloxy; R3 = (un) substituted benzyl; R4 = Ph, AB chloro-, bromo-, or fluorophenyl, (un)substituted benzyl] were prepd. and they inhibited aldose reductase. 2-Methyl-1-oxo-3-indanacetic acid was treated with BuONO, the I (R = Me, R1 = H, R2 = OH) obtained was dehydroxylated, and the product treated with 3,4-Cl2C6H3CH2Cl to give I (R = Me, R1 = H, R2 = 3,4-C12C6H3CH2).
- IT 79456-23-8P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 79456-23-8 HCAPLUS RN

4-Isoquinolineacetic acid, 1-[[(4-chlorophenyl)methyl)methylamino]-3-CN methyl- (9CI) (CA INDEX NAME)

III

IT 79456-22-7P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of and inhibitioin of aldose reductase by) 79456-22-7 HCAPLUS

RN

4-Isoquinolineacetic acid, 1-[[(4-chlorophenyl)methyl]methylamino)-3-methyl-, monohydrochloride (9CI) (CA INDEX NAME)

HC1

L30 ANSWER 21 OF 35 HCAPLUS COPYRIGHT 2001 ACS AN 1981:174912 HCAPLUS

94:174912 DN

ΤI Aminoisoquinoline derivatives

E. Gy. T. Gyogyszervegyeszeti Gyar, Hung. Neth. Appl., 20 pp. CODEN: NAXXAN PA

so

DT Patent

Dutch LA

FAN	CNT 1					
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
ΡI	NL 8002119	A	19801014	NL 1980-2119	19800411 <	
	ни 20959	0	19810928	HU 1979-EE2647	19790411 <	
	ни 178522	P	19820528			
	GB 2048256	A	19801210	.GB 1980-10793	19800331 <	
	GB 2048256	B2	19830518			
	BE 882674	A1	19801008	BE 1980-9778	19800408 <	
	AU 8057302	Al	19801016	AU 1980-57302	19800410 <	

ANSWER 21 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30-AN DN Aminoisoquinoline derivatives TI E. Gy. T. Gyogyszervegyeszeti Gyar, Hung. Neth. Appl., 20 pp.

CODEN: NAXXAN PA so DТ LA Dutch FAN. CNT 1 PATENT NO. KIND DATE PΙ APPLICATION NO. NL 8002119 HU 20959 DATE 19801014 NL 1980~2119 HU 178522 19810928 19800411 <--HU 1979-EE2647 GB 2048256 19820528 19790411 <--GB 2048256 A 19801210 GB 1980-10793 B2 BE 882674 19830518 19801008 1980033i <--AU 8057302 AI BE 1980-9778 Al 19801016 19800408 <--AU 1980-57302 19800410 <--

SEARCHED BY SUSAN HANLEY Phone: 305-4053

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AU 535462
                        В2
                              19840322
     FR 2453855
                        A1
                              19801107
                                              FR 1980-8052
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     FR 2453855
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                        В1
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                        С
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                                              DE 1980-3013998
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     DE 3013998
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                                                                19800411 <--
     ES 490506
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     CS 216218
                        B<sub>2</sub>
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                              19830730
                                                                19800411 <--
     AT 8001972
                        Α
                              19831015
                                              AT 1980-1972
                                                                19800411 <--
     AT 374798
                        В
                              19840525
PRAI HU 1979-EE2647
                              19790411
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I

ΑB Aminoisoquinolines I (R, R1 = H, alkyl; R2 = H, alkyl, optionally substituted Ph, pyridyl, dialkylaminoalkyl; NR1R2 = heterocyclic; R3 = alkoxy, amino) were prepd. Thus I (R = Me, R1 = R2 = H, R3 = Br) was treated with morpholine to give 83% I (R = Me, R1 = R2 = H, R3 = morpholino) which had a spontaneous motility-inhibiting ED50 of 400 mg/kg orally in mice. I (R-R2 = H, R3 = morpholino) had an analgesic ED50 of 100 mg/kg orally in the HOAc writhing test in mice and a therapeutic index

77454-38-7P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 77454-38-7 HCAPLUS RN

Isoquinoline, 1,3-di-4-morpholinyl- (9CI) (CA INDEX NAME) CN

ANSWER 22 OF 35 HCAPLUS COPYRIGHT 2001 ACS L30

1980:22393 HCAPLUS AN

DN 92:22393

1-Amino-4-phenylisoquinoline derivatives ΤI

Simmonds, Robin George TN

Aspro-Nicholas Ltd., Engl. PA

Brit., 16 pp. CODEN: BRXXAA so

DΥ Patent

LA English FAN. CNT 1

ΡI

GΙ

PATENT NO. KIND DATE APPLICATION NO. DATE GB 1545767 19790516 GB 1975-31144 19760630 <-- - - .

AB The prepn. is described of title compds. I (R, R1 = H, C1-12 alkyl; RNR1 = piperazinyl optionally substituted by C1-12 alkyl or hydroxyalkyl; n=0 piperazinyi optionally substituted by C1-12 alkyl or hydroxyalkyl; n = 0 - 3; m = 0 - 4; R2,R3 = C1-12 alkyl optionally substituted by .gtoreq.1 halo, C1-12 alkoxy, halo; R4 = H, C1-12 alkyl; R5,R6 = H or C1-12 alkyl, alkylthio, alkoxy; R5R6 = bond, O, S, C1-3 alkylene optionally contg. .gtoreq.1 O or S), which show antiinflammatory (esp. antirheumatic) and/or central nervous system activity. Thus, 3-dimethylamino-7,8-dihydrobenzo[1,2]cyclohepta[3.4.5-de]isoquinoline hydrogen maleate was prepd. from dibenzo[ad]suberone by sequential treatment with NaH/Me3S+ I-, BF3.Me2O/CH2Cl2, and H2NCO2Et/H2SO4 followed by heating (256.degree., 1 h), refluxing with POCl3, and Me2NH/EtOH treatment. The yields of the 6 steps were 96, 98, 100, 89, 99, and 75.6%, resp. Compns. contg. I are described.

TT 72240-39-2P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 72240-39-2 HCAPLUS

1-Isoquinolinamine, N,N,3-trimethyl-4-phenyl- (9CI) (CA INDEX NAME)

ANSWER 23 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1979:466274 HCAPLUS

DN 91:66274

Photographic products and processes employing nondiffusible ΤI 1-arylazo-4-isoquinolinol dye-releasing compounds

ΤN Chapman, Derek D.; Friday, James A.; Elwood, James K.

Eastman Kodak Co., USA PΑ

U.S., 21 pp. CODEN: USXXAM so

DT

Patent

English LA

FAN CNT 1

	L MIA .	CHIL					
PATENT NO.		KIND DATE	DATE	APPLICATION NO.	DATE		
	PI	US 4148642	A	19790410	US 1978-884469	19780307 <	
		US 4183754	A	19800115	US 1978-950194	19781010 <	
	PRAI	US 1978-884469		19780307			
	GT						

CONH (CH₂) 40 C5H₁₁-tert

$$C5H_{11}$$
-tert

OH

N=N

OH

AB Photog. elements, diffusion-transfer assemblages, and processes are described which employ a novel nondiffusible compd. having a releaseable 1-arylazo-4-isoquinolinol dye moiety. The compd. contains in the ortho position of the arylazo moiety a metal chelating group, a salt thereof, or a hydrolyzable precursor thereof, and a ballasted carrier moiety which is capable of releasing the diffusable azo dye under alk. conditions. dye is transferred imagewise to an image-receiving layer where it is contacted with metal ions to form a metal complexed azo dye transfer image of excellent stability. Thus, a single-color integral-imaging receiver element was prepd. by coating successively on a polyester film support a metalizing layer comprising gelatin (1.08 g/m2) and NiSo4.6H2O (0.58 g/m2), a receiving layer consisting of gelatin and poly(4-vinylpyridine) (each at 2.15 g/m2), a reflecting layer comprising TiO2 and gelatin in 6.25/1 ratio, an opaque layer of C in gelatin, a layer consisting of gelatin and a dispersion of I (prepd. by reaction of 3-methyl-4isoquinolinol with diazotized 4-(3-amino-4-hydroxybenzenesulfonamido)-1hydroxy-N-[4-(2,4-di-tert-phenylphnoxybutyl]-2-naphthamide) (0.84 g/m2), a layer of red sensitized internal image emulsion, a layer of dodecylhydroquinone(1.29 g/m2) dispersed in gelatin (1.61 g/m2), and a gelatin overcoat layer. This integral element was exposed to a multicolor test object and then processed to show a d. at .lambda.max after 4 min of 1.11, a .lambda.max of 637 nm, a half bandwidth of 127, and a d. change after exposure to a 5000 ft-candle light source for 2 days of -0.03 vs 1.41, 544, 186, and -0.18, resp., for a Ni2+ -free control. 70881-90-2

RL: USES (Uses)

(azo dye-releasing compd., for color photog.) 70881-90-2 HCAPLUS

 $\begin{tabular}{ll} 2-Naphthalenecarboxamide, & N-[4-[2,4-bis(1,1-dimethylpropyl)phenoxy]butyl]-1-hydroxy-4-[[4-hydroxy-3-methyl-1-meth$ isoquinolinyl)azo]phenyl]sulfonyl]amino]- (9CI) (CA INDEX NAME)

RN 70881-92-4 HCAPLUS
CN Benzoic acid, 2-{(4-hydroxy-3-methyl-1-isoquinolinyl)azo}- (9CI) (CA INDEX NAME)

RN 70881-93-5 HCAPLUS
CN Benzenesulfonamide, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo](9CI) (CA INDEX NAME)

RN 70881-94-6 HCAPLUS
CN 4-Isoquinolinol, 1-[[2-hydroxy-4-(methylsulfonyl)phenyl]azo]-3-methyl(9CI) (CA INDEX NAME)

RN 70881-95-7 HCAPLUS
CN 4-Isoquinolinol, 1-[[2-hydroxy-5-[(trifluoromethyl)sulfonyl]phenyl]azo]-3methyl- (9CI) (CA INDEX NAME)

RN 70881-96-8 HCAPLUS

CN Benzoic acid, 2-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-5-nitro- (9CI) (CA INDEX NAME)

RN 70881-97-9 HCAPLUS

CN 4-Isoquinolinol, 1-[(2-hydroxy-4-nitrophenyl)azo]-3-methyl- (9CI) (CA INDEX NAME)

RN 70882-04-1 HCAPLUS

CN Benzenesulfonamide, 4-hydroxy-3-[(4-hydroxy-3-methyl-1-isoquinolinyl)azo]-N-(3-hydroxyphenyl)- (9CI) (CA INDEX NAME)

ANSWER 24 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1977:601283 HCAPLUS

DN 87:201283

TI Synthesis and antitussive activity of 3-azabicyclo[3.2.2]nonane derivatives

ΑU

CS

Arya, V. P.; Kaul, C. L.; Grewal, R. S. Ciba-Geigy Res. Cent., Bombay, India Arzneim.-Forsch. (1977), 27(9), 1648-52 SO CODEN: ARZNAD

DΤ Journal

LA English

GI

AB Mannich bases I (R = Me, H, Et, Pr; Rl = 4-FC6H4, 4-PhCH2OC6H4, 4-BrC6H4, 4-ClC6H4, 3-pyridyl, 3-indolyl, 2-thienyl), prepd. from the substituted acetophenones and propiophenones and 3-azabicyclo[3.2.2]nonane, were evaluated for antitussive activity. I (R = Me, Rl = 4-PhCH2OC6H4) (II) was as potent as codeine and dextromethorphan in its antitussive activity. II also exhibited antimorphine activity. There was no direct correlation between the antitussive effect and antimorphine activity.

ΙT 64686-73-3P RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 64686-73-3 HCAPLUS RN

3-Azabicyclo[3.2.2]nonane, 3-[3-[(3-azabicyclo[3.2.2]non-3-y1)methy1]-4methyl-1-isoquinolinyl]-, trihydrochloride (9CI) (CA INDEX NAME)

19750425 <--

19790214 <--

3 HC1

```
ANSWER 25 OF 35 HCAPLUS COPYRIGHT 2001 ACS
AN
     1976:560167 HCAPLUS
DN
     85:160167
ΤŢ
     Piperazinoisoquinolines
PA
     Thomae, Dr. Karl, G.m.b.H., Ger.
     Fr. Demande, 22 pp.
SO
     CODEN: FRXXBL
DΤ
     Patent
LA
     French
FAN.CNT 2
     PATENT NO.
                      KIND
                            DATE
                                            APPLICATION NO.
    FR 2268524
                       A1
                            19751121
                                            FR 1975-13095
     DE 2420012
                       Al
                            19751120
                                            DE 1974-2420012 19740425 <--
     DE 2420012
                       B2
                            19790517
    DE 2420012
                       Ç3
                            19800110
    DE 2503961
                            19760805
                                            DE 1975-2503961 19750131 <--
    DE 2503961
                            19790705
    DE 2503961
                       C3
                            19800228
```

19800115

19740425

19750131

19750423

I

СН 615180

PRAI DE 1974-2420012

CH 1975-5155

DE 1975-2503961

- Piperazinylisoquinolines I (R = H, R1 = H, 5-Me, 5-Cl, 7-Cl, 5-F, 5-OMe, 5-NO2, X = S, SO; R = Ac, CHO, R1 = H, X = SO; R = Ac, CO2Et, Me, H, R1 = 5-Me, X = SO; R = H, Me, R1 = H, X = O; R = Ac, R1 = 5-NO2, X = S) were prepared. e. g. by treating 1,3-dichloroisoquinoline with the morpholine derive followed by treating with a piperarise derive follower by the followed by the followed by the followed by the follower by the follow AB deriv. followed by treating with a piperazine deriv. I are platelet aggregation inhibitors. Thus in the test according to Morris I (R = H, R1 = 5-C1, X = S0) gave 92% inhibition at 10-4 mole/1.
- IT 60691-16~9P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and acylation of) 60691-16-9 HCAPLUS
- Isoquinoline, 5-nitro-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, monohydrochloride (9CI) (CA INDEX NAME)

CH 1979-1429

```
ANSWER 25 OF 35 HCAPLUS COPYRIGHT 2001 ACS
      1976:560167 HCAPLUS
85:160167
AN
DN
      Piperazinoisoquinolines
      Thomae, Dr. Karl, G.m.b.H., Ger. Fr. Demande, 22 pp.
SO
      CODEN: FRXXBL
DT
      Patent
      French
FAN.CNT 2
      PATENT NO.
                                                  APPLICATION NO. DATE
                          KIND DATE
                                                                      19750425 <--
PΙ
      FR 2268524
                                 19751121
                                                  FR 1975-13095
      DE 2420012
                                                  DE 1974-2420012 19740425 <--
                           Al
                                 19751120
                                 19790517
      DE 2420012
      DE 2420012
DE 2503961
                          C3
                                 19800110
                                 19760805
                                                  DE 1975-2503961 19750131 <--
      DE 2503961
DE 2503961
                                 19790705
                                 19800228
                           C3
      CH 615180
                                 19800115
                                                  CH 1979-1429
                                                                      19790214 <--
PRAI DE 1974-2420012
DE 1975-2503961
                                 19740425
19750131
      CH 1975-5155
                                 19750423
GI
```



Piperazinylisoquinolines I (R = H, Rl = H, 5-Me, 5-Cl, 7-Cl, 5-F, 5-OMe, 5-NO2, X = S, SO: R = Ac, CHO. Rl = H, X = SO: R = Ac, CO2Et. Me, H, Rl = 5-Me, X = SO: R = H, Me, Rl = H, X = O: R = Ac, Rl = 5-NO2, X = S) were prepd. e. g. by treating 1,3-dichloroisoquinoline with the morpholine deriv. followed by treating with a piperazine deriv. I are platelet aggregation inhibitors. Thus in the test according to Morris I (R = H, Rl = 5-Cl, X = SO) gave 92% inhibition at 10-4 mole/1.

60691-16-9P IT

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and acylation of) 60691-16-9 HCAPLUS

Isoquinoline, 5-nitro-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, monohydrochloride (9CI) (CA INDEX NAME)

● HCl

IT 58138-21-9P 60691-07-8P 60691-10-3P 60691-12-5P 60691-13-6P 60691-15-8P 60691-17-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)

(prepn. and oxidn. of)
RN 58138-21-9 HCAPLUS

CN Isoquinoline, 3-(1-piperazinyl)-1-(4-thiomorpholinyl)- (9CI) (CA INDEX NAME)

RN 60691-07-8 HCAPLUS

CN Isoquinoline, 5-methyl-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, monohydrochloride (9CI) (CA INDEX NAME)

HC1

RN 60691-10-3 HCAPLUS

CN Isoquinoline, 5-chloro-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, monohydrochloride (9CI) (CA INDEX NAME)

HC1

RN 60691-12-5 HCAPLUS
CN Isoquinoline, 7-chloro-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-,
(2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM

CRN 60691-11-4 CMF C17 H21 C1 N4 S

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

RN 60691-13-6 HCAPLUS
CN Isoquinoline, 5-fluoro-3-(1-piperazinyl)-1-(4-thiomorpholinyl)~ (9CI) (CA INDEX NAME)

RN

60691-15-8 HCAPLUS
Isoquinoline, 5-methoxy-3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, sulfate (2:1) (9CI) (CA INDEX NAME) CN

CM 1

CRN 60691-14-7 CMF C18 H24 N4 O S

CM

CRN 7664-93-9 CMF H2 O4 S

RN

60691-17-0 HCAPLUS
Piperazine, 1-acetyl-4-(5-nitro-1-(4-thiomorpholinyl)-3-isoquinolinyl}-(9CI) (CA INDEX NAME) CN

58138-22-0P 58138-25-3P 60691-09-0P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. and platelet aggregation inhibiting activity of) 58138-22-0 HCAPLUS Isoquinoline, 3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, (22)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM

RN CN

> CRN 58138-21-9 CMF C17 H22 N4 S

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

RN 58138-25-3 HCAPLUS
CN Isoquinoline, 1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-,
(2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM

CRN 58138-24-2 CMF C17 H22 N4 O S

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z Double bond geometry as shown.

60691-09-0 HCAPLUS RN

Isoquinoline, 3-(4-methyl-1-piperazinyl)-1-(4-morpholinyl)- (9CI) (CA CN INDEX NAME)

IT

60691-18-1P 60691-19-2P 60691-22-7P RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. and platelet aggregation-inhibiting activity of) 60691-18-1 HCAPLUS

· RN

Isoquinoline, 5-methyl-1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-CN (9C1) (CA INDEX NAME)

RN 60691-19-2 HCAPLUS

Isoquinoline, 5-chloro-1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-CN (9CI) (CA INDEX NAME)

RN 60691-22-7 HCAPLUS

Isoquinoline, 5-methoxy-1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-, CN monohydrochloride (9CI) (CA INDEX NAME)

HC1

• HC1

RN 58138-24-2 HCAPLUS
CN Isoquinoline, 1-(1-oxido-4-thiomorpholiny1)-3-(1-piperaziny1)- (9CI) (CA INDEX NAME)

RN 60691-08-9 HCAPLUS
CN Isoquinoline, 1-(4-morpholinyl)-3-(1-piperazinyl)-, monohydrochloride

(9CI) (CA INDEX NAME)

● HC1

RN 60691-20-5 HCAPLUS
CN Isoquinoline, 7-chloro-1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-(9CI) (CA INDEX NAME)

RN 60691-21-6 HCAPLUS
CN Isoquinoline, 5-fluoro-1-(1-oxido-4-thiomorpholiny1)-3-(1-piperaziny1)(9CI) (CA INDEX NAME)

RN 60691-24-9 HCAPLUS
CN Isoquinoline, 5-nitro-1-(1-oxido-4-thiomorpholinyl)-3-(1-piperazinyl)-,
(2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM :

CRN 60691-23-8 CMF C17 H21 N5 O3 S

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

RN 60691-25-0 HCAPLUS
CN Piperazine, 1-acetyl-4-[1-(1-oxido-4-thiomorpholinyl)-3-isoquinolinyl)(9CI) (CA INDEX NAME)

RN 60691-26-1 HCAPLUS
CN 1-Piperazinecarboxaldehyde, 4-[1-(4-morpholinyl)-3-isoquinolinyl]- (9CI)
(CA INDEX NAME)

RN 60691-27-2 HCAPLUS
CN Piperazine, 1-acetyl-4-[5-methyl-1-(1-oxido-4-thiomorpholinyl)-3isoquinolinyl]- (9CI) (CA INDEX NAME)

RN 60691-28-3 HCAPLUS

1-Piperazinecarboxylic acid, 4-[5-methyl-1-(1-oxido-4-thiomorpholinyl)-3-isoquinolinyl]-, ethyl ester (9CI) (CA INDEX NAME) CN

60691-29-4 HCAPLUS RN

Isoquinoline, 5-methyl-3-(4-methyl-1-piperazinyl)-1-(1-oxido-4thiomorpholinyl)- (9CI) (CA INDEX NAME) CN

ANSWER 26 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1976:59571 HCAPLUS L30

AN

DN 84:59571

TI Isoquinolines

Nickl, Josef; Mueller, Erich; Schroeter, Wolfgang; Haarmann, Walter Thomae, Dr. Karl, G.m.b.H., Ger. In

Ger. Offen., 18 pp. CODEN: GWXXBX **\$**0

DT Patent

German LA

	PATENT NO.	KIND	DATE	APE	LICATION NO.	DATE	•
I	DE 2420012	A1	19751120	DE	1974-2420012	19740425	<
	DE 2420012	B2	19790517				
	DE 2420012	C3	19800110				
	AT 7501639	A	19770715	AT	1975-1639	19750304	<
	NL 7504016	A	19751028	NL	1975-4016	19750404	<
	ES 436317	A1	19770201	ES	1975-436317	19750404	<~~
	FI 7501067	Α	19751026	FI	1975-1067	19750409	<
	FI 61882 '	В	19820630				
	FI 61882	С	19821011				
	DK 7501579	A	19751026	DK	1975-1579	19750411	<
	DK 140841	В	19791126	•			
	DK 140841	С	19800505				
	US 3975524	A	19760817	US	1975-567234	19750411	<
	SU 557756	D	19770505	SU	1975-2121918	19750411	<
	DD 119047	С	19760405	DD	1975-185646	19750423	<
	RO 66020	В	19790815	RO	1975-82052	19750423	<
	RO 66020	P	19800615				
	CH 613965	A	19791031	CH	1975-5155	19750423	<
	BE 828355	A1	19751024	BE	1975-155746	19750424	<
	NO 7501473	Α	19751028		1975-1473	19750424	
	NO 142403	В	19800505				
	NO 142403	č	19800813				
	JP 50142578	A2	19751117	JР	1975-50160	19750424	<
	JP 58004020	B4	19830124	V-	15.5 50100	13.30121	-
	AU 7580511	Al	19761028	AII	1975-80511	19750424	<
	ZA 7502649	A	19761229		1975-2649	19750424	
	GB 1466227	A	19770302		1975-17085	19750424	
	HU 170231	P	19770428		1975-TO1001	19750424	
	PL 93821	p	19770630		1975-179891	19750424	
	IL 47155	A1	19780310		1975-47155	19750424	
	SE 404926	В	19781106		1975-4779	19750424	
	SE 404926	Č	19790215		13/3 1//3	13750124	•
	CA 1051893	A1	19790403	CA	1975-225579	19750424	<i><</i>
	FR 2268524	A1	19751121		1975-13095	19750425	
	CS 193512	P	19791031		1975-2918	19750425	
	ES 439038	A1	19770201		1975-439038	19750701	
λт	DE 1974-2420012	N1	19740425	ES	1973-439030	13/30/01	
wı	DE 1974-2403961		19750131				•
	DE 1975-2503961		19750131				
}	For diagram(s), Isoquinolines I				troating 1 2-	dichloroi	inol
,	with thiomorphol	(n = 0	ito S-ovido	pu. by	creating 1,3-	42 - 07	aodatuor
	oxidized to I (n	1116 OT	Tre proxime	ve 3 + rd birp	imas 10-5 ms	11 - U) w	as also
	thrombocyte aggr						
		egraci	on in the th	rompocy	te stickiness	cesc by	re and a
,	resp.						
•	58138-24-2P	ia ===	namation) - D	DED /D-	onamation)		
	RL: SPN (Synthet						
			cyce aggrega	cion in	hibiting acti	ATEN OF)	
!	58138-24-2 HCAP			-1-11			.0071
ı	Isoquinoline, 1- INDEX NAME)	(1-0X1	do-4-thiomor	Биотти?	T)-3-(I-biber	azınyı)-	(9CI) (

CRN 58138-21-9

CMF C17 H22 N4 S

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

RN 58138-23-1 HCAPLUS
CN Isoquinoline, 3-(1-piperazinyl)-1-(4-thiomorpholinyl)-, monohydrochloride
(9CI) (CA INDEX NAME)

● HCl

CM

CRN 58138-24-2 CMF C17 H22 N4 O S

CM

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

ΙT 58138-21-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn., oxidn., and thrombocyte aggregation inhibiting activity of) 58138-21-9 HCAPLUS

Isoquinoline, 3-(1-piperazinyl)-1-(4-thiomorpholinyl)- (9CI) (CA INDEX

L30 ANSWER 27 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1975:16836 HCAPLUS AN

DN 82:16836

Hypolipemic and hypoglycemic 1-(1-imidazoly1)isoquinolines Lerch, Ulrich; Granzer, Ernold TI

IN

Farbwerke Hoechst A.-G. PA

Ger. Offen., 34 pp. CODEN: GWXXBX so

DT Patent

LA German

FAN.CNT 1					
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	DE 2314985	A1	19741017	DE 1973-2314985	19730326 <
	ES 424436	A1	19761101	ES 1974-424436	19740320 <
	GB 1464289	A	19770209	GB 1974-12861	19740322 <
	ZA 7401917	Α	19750326	ZA 1974-1917	19740325 <
	DD 114607	Ç	19750812	DD 1974-177438	19740325 <

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AU 7467098
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                                                                      19740325 <--
      US 3914236
                                 19751021
                                                  US 1974-454713
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      HU 168524
                                 19760528
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                                                                      19740325 <--
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                                 19761015
                                                  AT 1974-2452
                           Α
                                                                      19740325 <--
      AT 337183
                                 19770610
                           В
                                 19740926
      BE 812841
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                           A1
                                                                      19740326 <--
      FR 2223024
                           A1
                                 19741025
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                                                                      19740326 <--
      JP 49126684
                           A2
                                 19741204
                                                  JP 1974-33183
                                                                      19740326 <--
      US 3961062
                                 19760601
                                                  US 1975-562048
                                                                      19750326 <--
PRAI DE 1973-2314985
                                 19730326
      DE 1973-7314985
                                 19730326
      US 1974-454713
                                 19740325
      For diagram(s), see printed CA Issue.
     Nineteen imidazolyl-isoquinolines I (R=H, Cl, Ph, or Et; Rl=H, Ph, cyclohexyl, Et, Bu, or Cl; R2, R3, R4=H, Ph, or Me) and (or) their
      salts, e.g. hydrochlorides, were prepd. by reaction of the corresponding 1-chloroisoquinolines with the imidazoles in the presence of NaH or KOH or
      Bu3N in, e.g., (MeOCH2)2 or DMF. I had hypolipemic and hypoglycemic
      activities in rats and rabbits.
ΙT
      55150-98-6P 55151-06-9P 55151-07-0P
      55151-08-1P 55151-09-2P
      RL: SPN (Synthetic preparation); PREP (Preparation)
      (prepn. and hypoglycemic and hypolipemic activity of) 55150-98-6 HCAPLUS
RN
      Isoquinoline, 4-chloro-1-(1H-imidazol-1-yl)-3-phenyl- (9CI) (CA INDEX
CN
```

● HCl

RN 55151-07-0 HCAPLUS
CN Isoquinoline, 4-chloro-3-ethyl-1-(1H-imidazol-1-yl)-, phosphate (9CI) (CA INDEX NAME)

CM 1

CRN 55150-96-4 CMF C14 H12 C1 N3

200928-22-9 HCAPLUS RN 1,7-Naphthyridine, 8-(1-methylhydrazino)-6-(3-methyl-2-pyridinyl)- (9CI) CN (CA INDEX NAME)

RN 200928-24-1 HCAPLUS Glycine, N-[[4-(acetylamino)phenyl]sulfonyl]-N-[6-(3-methyl-2-pyridinyl)-n-[6-(3-methyl-2-pyridinyl)-n-[6-(3-methyl-2-pyridinyl)]CN 1,7-naphthyridin-8-yl]-, ethyl ester (9CI) (CA INDEX NAME)

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ANSWER 2 OF 35 HCAPLUS COPYRIGHT 2001 ACS
L30
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AN 1997:498744 HCAPLUS

DN 127:190707

Synthesis and antitumor activity of 3-arylisoquinoline derivatives TI

Cho, Won-Jea: Yoo, Su-Jeong: Park, Myun-Ji; Chung, Byung-Ho; Lee, ΑU Chong-Ock

College of Pharmacy, Chonnam National University, Kwangju, 500-757, S. cs

Korea

Arch. Pharmacal Res. (1997), 20(3), 264-268 CODEN: APHRDQ; ISSN: 0253-6269 Pharmaceutical Society of Korea SO

PΒ DT Journal

English

GΙ

CM

CRN 7664-38-2 CMF H3 O4 P

55151-08-1 HCAPLUS RN Isoquinoline, 1-(1H-imida2ol-1-yl)-3-phenyl-, ethanedioate (9CI) (CA INDEX NAME)

СМ

CRN .55150-97-5 CMF C18 H13 N3

CM

CRN 144-62-7 CMF C2 H2 O4

RN CN

55151-09-2 HCAPLUS
Isoquinoline, 4-chloro-1-(lH-imidazol-1-yl)-3-phenyl-, ethanedioate (9CI)
(CA INDEX NAME)

CM 1

CRN 55150-98-6 CMF C18 H12 C1 N3

CM

CRN 144-62-7 CMF C2 H2 O4

55150-95-3P 55150-96-4P 55150-97-5P IT RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of hypoglycemic and hypolipemic) 55150-95-3 HCAPLUS

Isoquinoline, 3-ethyl-1-(lH-imidazol-1-yl)- (9CI) (CA INDEX NAME) CN

55150-96-4 HCAPLUS

Isoquinoline, 4-chloro-3-ethyl-1-(1H-imidazol-1-yl)- (9CI) (CA INDEX NAME)

RN 55150-97-5 HCAPLUS

Isoquinoline, 1-(1H-imidazol-1-yl)-3-phenyl- (9CI) (CA INDEX NAME) CN

L30 ANSWER 28 OF 35 HCAPLUS COPYRIGHT 2001 ACS AN 1972:564414 HCAPLUS DN 77:164414 Reactions of 1-chloro-3-chloromethyl-4-methylisoquinoline ΤI AU Nair, M. D. CS Ciba Res. Cent., Bombay, India Indian J. Chem. (1972), 10(4), 337-40 SO CODEN: IJOCAP DT Journal LA English GI For diagram(s), see printed CA Issue. with secondary bases 1-chloro-3-(chloromethyl)-4-methylisoquinoline (I) gave mono or disubstitution products in which the Cl in positions 1 or 3, or both was replaced. In 1-chloro-3-[(2-methylpiperidino)-methyl]-4-methylisoquinoline there was NMR evidence for non-equivalence of benzylic methylene protons from the asymmetry of the 2-Me substituent on piperidine. Reaction of I with piper-azine gave a bis condensation product, II, with NH3 and 4-(.gamma.-aminopropyl)morpholine III and IV were obtained, resp. Nitra-tion of I gave the corresponding 5-NO2 deriv., reaction of which with bases gave mono or disubstituted products, depending on reaction conditions. 14576-16-0P 14576-17-1P 14577-67-4P 14657-46-6P 14657-48-8P 14657-49-9P 14657-50-2P 14657-51-3P 14657-52-4P 18716-17-1P 37978-50-0P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) 14576-16-0 HCAPLUS 1-Piperazineethanol, 4-[[1-[4-(2-hydroxyethy1)-1-piperaziny1]-4-methy1-3-isoquinoliny1]methy1]- (9CI) (CA INDEX NAME)

RN 14576-17-1 HCAPLUS
CN 1-Piperazinecarboxylic acid, 4-[{1-[4-(ethoxycarbonyl)-1-piperazinyl]-4methyl-3-isoquinolinyl}methyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN

14577-67-4 HCAPLUS
Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]- (8CI, 9CI) (CA INDEX NAME) CN

RN 14657-46-6 HCAPLUS

CN Isoquinoline, 4-methyl-1-(1-piperidinyl)-3-(1-piperidinylmethyl)- (9CI) (CA INDEX NAME)

RN

14657-48-8 HCAPLUS
Isoquinoline, 4-methyl-3-((2-methyl-1-piperidinyl)methyl]-1-(4-morpholinyl)- (9CI) (CA INDEX NAME) CN

RN 14657-49-9 HCAPLUS
CN Isoquinoline, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-5-nitro(9CI) (CA INDEX NAME)

RN 14657-50-2 HCAPLUS
CN Isoquinoline, 4-methyl-5-nitro-1-(1-piperidinyl)-3-(1-piperidinylmethyl)(9CI) (CA INDEX NAME)

RN 14657-51-3 HCAPLUS
CN 5-Isoquinolinamine, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)(9CI) (CA INDEX NAME)

RN 14657-52-4 HCAPLUS CN Isoquinoline, 5-chloro-4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)- (9CI) (CA INDEX NAME)

RN 18716-17-1 HCAPLUS
CN Isoquinoline, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-,
(2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 18704-39-7 CMF C19 H25 N3 O2

CM 2

CRN 110-16-7 CMF C4 H4 O4 CDES 2:Z

Double bond geometry as shown.

RN 37978-50-0 HCAPLUS
CN Isoquinoline, 4-methyl-1-(1-pyrrolidinyl)-3-(1-pyrrolidinylmethyl)-, trihydrochloride (9CI) (CA INDEX NAME)

●3 HCl

ANSWER 29 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1971:544847 HCAPLUS 75:144847 Crystal structure of 5-hydroxy-3-phenyl-1-(3-methyl-1-isoquinolyl)pyrazole TI King, Geoffrey S. D.; Reimlinger, Hans ΑU Union Carbide Eur. Res. Assoc., Brussels, Belg. CS Chem. Ber. (1971), 104(9), 2694-701 CODEN: CHBEAM DT Journal LA German An x-ray crystal structure detn. of the title compd. (I) proved that I is the product of the reaction of PhC.tplbond.CCO2Me with 1-hydrazino-3-methylisoquinoline. I crystd. orthorhombic with a 43.26, b 12.626, c 5.546 .ANG., d.(exptl.) 1.32, d.(calcd.) 1.321, and space group P212121, and the asym. unit contained 2 independent mols. 34274-79-8 RL: PRP (Properties) (crystal structure of) 34274-79-8 HCAPLUS RN

Pyrazol-5-ol, 1-(3-methyl-1-isoquinolyl)-3-phenyl- (8CI) (CA INDEX NAME)

- HO N N
- ANSWER 30 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1970:425363 HCAPLUS DN Condensed isoquinolines. I. Syntheses of s-triazolo[3,4-a]isoquinolines Reimlinger, Hans; Vandewalle, Jan J. M.; Lingier, Willy R. F. ŤΙ AU Union Carbide European Res. Assoc., Brussels, Belg. CS Chem. Ber. (1970), 103(6), 1960-81 SO CODEN: CHBEAM DT Journal LA German For diagram(s), see printed CA Issue. GI Hexasubstituted s-triazolo[3,4-a]isoquinolines(I) [where R = H, Me, CH2CO2Et, CH2NHBz, CH2CH2CL, CO2Et, CF3, CH2CH2NHBz, o-ClC6H4, Et, CH2CH2CO2H, Ph, HC:CPh, CH:CHPh.HCl, 3-indolylmethyl, CH2C6H3(OMe)2-3,4, n-C17H35, CH2CONHEt, CH2CONHMe, CH2CONMe2, CH2CH2OH, CH2CO2H, CHPh2, NHPh, cyclohexylamino, 1-pyridyl, or 4-pyridyl; Rl = H or Cl; R2, R4 = H or MeO;

PATEL 09/852,850

R3 = H or NO2; and R5 = H, Cl, or OMe] were prepd. by 1 or more of several methods: (a) by reaction of 1,4-dichloroisoquinoline (II) with N2H4 and RCO2Bu, (b) treatment of 1-hydrazinoisoquinoline with RCOCl, (c) reaction of II with NH2NHCOR, or (d) treatment of 1-[2-(RCOsubstituted)hydrazino)isoquinoline with SOC12. Reaction of 1-hydrazino-3,4-(RR1-disubstituted) isoquinolines (III) with Cl2C:X yielded disubstituted 1,2-dihydro-s-triazolo(3,4-a)isoquinolines (IV) (where R = H, Cl, or Me, Rl = H or Cl, and X = O, S, NH, or NBz). 27319-97-7P RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 27319-97-7 HCAPLUS

3-Pyrazolin-5-one, 1-(3-methyl-1-isoquinolyl)-3-phenyl- (8CI) (CA INDEX

IΤ

L30 ANSWER 31 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1968:435972 HCAPLUS AN

69:35972 DN

4-Methylisoquinolines TI

Aebi, Albert; Nair, Mohan D.; Bucher, Karl IN

PA CIBA Ltd.

Swiss, 6 pp CODEN: SWXXAS

DT Patent

LA German

FAN.CNT 1

PATENT NO. APPLICATION NO. KIND DATE DATE --------------

CH 438308 19671215 CH PΤ GT

For diagram(s), see printed CA Issue. The title compds. are prepd. by treating 1-chloro-3-chloromethyl-4methylisoquinoline (I) or its substituted derivs. with secondary amines. Thus, 1.55 g. I and 5 ml. morpholine was heated overnight in a pressure vessel at 150.degree.. The cryst. suspension was then evapd. to dryness, taken up in CHCl3, extd. 2 times with dil. aq. HCl, and the aq. exts. adjusted to pH 8-9 with NaOH. The oil which sepd. gradually crystd., and was sepd. and recrystd. from iso-PrOH to give II (R = H and R1 = morpholino), m. 100.degree.; dihydrochloride m. 229-32.degree. (decompn.) and maleate m. 173-5.degree.. Other II similarly prepd. are shown in the table. The starting material for II (R = NO2) was prepd. by treating I with concd. H2SO4 and fuming HNO3 to give II (R = NO2, R1 = C1), m. 104-5.degree.. A mixt. of 4 g. 1,7-dichloro-3-chloromethyl-4methylisoquinoline (IV) and 50 ml. morpholine was refluxed 4 hrs., and excess morpholine was then removed under reduced pressure. [TABLE OMITTED] The residue was treated with aq. Na2CO3 until alk. and extd. with CHCl3. The exts. were evapd. to give 7-chloro-4-methyl-1-morpholino-3-(morpholinomethyl)isoquinoline, which was purified by conversion to its maleate and then to the free base, m. 120.degree. (EtOH). IV was prepd. by treating 4,4-dimethylhomophthalimide with fuming HNO3 and concd. H2SO4 at -10.degree. to give 4,4-dimethyl-7-nitrohomophthalimide, m. 209-11.degree.. Hydrogenation over Pd-C gave the 7-amino compd., m. 176-9.degree., which was diazotized and treated with CuCl to give the

7-chloro deriv., m. 200.degree.. Treatment with POC13 gave IV, m. 135.degree.. These compds. are used in pharmaceutical applications.

14576-16-0P 14576-17-1P 14577-67-4P

19630221 <--

RN 14576-17-1 HCAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 14577-67-4 HCAPLUS

CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]- (8CI, 9CI) (CA INDEX NAME)

RN 14657-46-6 HCAPLUS

N Isoquinoline, 4-methyl-1-(1-piperidinyl)-3-(1-piperidinylmethyl)- (9CI)

- -(CA INDEX NAME) - - -

RN 14657-49-9 HCAPLUS

CN Isoquinoline, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-5-nitro-(9CI) (CA INDEX NAME)

RN 14657-50-2 HCAPLUS

CN Isoquinoline, 4-methyl-5-nitro-1~(1-piperidinyl)-3-(1-piperidinylmethyl)(9CI) (CA INDEX NAME)

RN 14657-51-3 HCAPLUS

CN 5-Isoquinolinamine, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)(9CI) (CA INDEX NAME)

RN 14657-52-4 HCAPLUS

- CN Isoquinoline, 5-chloro-4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)- (9CI) (CA INDEX NAME)

- RN 14825-52-6 HCAPLUS
- CN 1-Piperazineethanol, 4,4'-{methylene(4-methyl-3,1-isoquinolinediyl)}di-, hydrochloride (8CI) (CA INDEX NAME)

•x HCl

- RN 18704-39-7 HCAPLUS
- CN Isoquinoline, 4-methyl-1-morpholino-3-(morpholinomethyl)- (8CI) (CA INDEX NAME)

- RN 18704-40-0 HCAPLUS
- CN Isoquinoline, 4-methyl-1-morpholino-3-(morpholinomethyl)-, dihydrochloride (8CI) (CA INDEX NAME)

●2 HC1

RN 18704-43-3 HCAPLUS
CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]-, monohydrochloride (8CI) (CA INDEX NAME)

HC1

RN 18716-17-1 HCAPLUS CN Isoquinoline, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-, (2Z)-2-butenedioate (1:1) (9CI) (CA INDEX NAME)

CM 1

.CRN 18704-39-7 CMF C19 H25 N3 O2

CM :

CRN 110-16-7 CMF C4 H4 O4 CDES .2: Z - - -

Double bond geometry as shown.

ANSWER 32 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1968:418996 HCAPLUS DN 69:18996 3-Chloroisocarbostyril and its chlorination products ΤI Nair, M. D.; Mehta, S. R. AU CS CIBA Res. Centre, Goregaon, India Indian J. Chem. (1967), 5(10), 467-70 SO CODEN: IJOCAP DT Journal English GΙ For diagram(s), see printed CA Issue. A mixt. of 10 g. dried homophthalimide (I) and 24 ml. POC13 was refluxed 14 hrs. (anhyd. conditions) at 170.degree., the soln. cooled and EtOH added to yield 8 g. 3-chlorocarbostyril (II), m. 219-20.degree. (EtOH). Methylation of I with MeI and alc. KOH soln. yielded 2-methyl-3chloroisocarbostyril (III), m. 109-10.degree. (EtOH). III was also prepd. by heating 1 hr. 15 g. N-methylhomophthalimide, 30 ml. POC13, and 1 ml. H2O at 170.degree. (oil-bath) and working up of the reaction mixt. A mixt. of 60 g. homophthalic acid and 100 ml. iso-PrNH2 and 20 ml. H2O was evapd. to dryness in vacuo, the residue mixed with 150 ml. o-Cl2C6H4 and heated overnight at 170.degree. to yield 52 g. N-isopropylhomophthalimide (IV), m. 88-9.degree. (EtOH). A mixt. of 8 g. IV, 24 ml. POCl3, and 1 ml. concd. HCl was heated 1 hr. at 170.degree. to yield 4 g. 1,3-dichloroisoquinoline, m. 121-2.degree. II was treated with a no. of amines to give 3-aminoisocarbostyrils (V). Thus, II was mixed with approx. 5 times its wt. of secondary amine, and the mixt. heated 8 hrs. at 150.degree. in a bomb tube to yield the following V (R = H) (R1, % yield, and m.p. given): morpholino, 39, 212.degree. (CHCl3-petroleum ether); pyrrolidino, 58, 238-41.degree. (decompn.); piperidino, 51.5, 195-7.degree. (CHCl3-petroleum ether); N-methylpiperazino, 47 212.degree.; hexamethylenimino, 59.5, 177-9.degree.; N-carbethoxypiperazino, 44.7, 196-7.degree.; 4-methylpiperidino, 55.7, 230-2.degree.; tetrahydroisoquinolino, 54.5, 217-18.degree.; N-(.beta.-hydroxyethyl)-piperazino (Va), 37.5, 205-7.degree.. Also prepd. was 19.8% V (R = Me, R1 = morpholino), m. 131-2.degree.. A mixt. of 2 g. 3-[N4-(.beta.-hydroxyethyl)piperazino)isocarbostyril (Va) and 10 ml. POC13 was refluxed 3 hrs. to yield 2.4 g. 1-chloro-3-[N4-(.beta.-chloroethyl)-piperazino)isoquinoline (VI) HCl salt, m. 300.degree. (EtOH-ether). VI (3.2 g.) on refluxing 18 hrs. with 15 ml. morpholine yielded \(\) 1-morpholino-3-(N4-(.beta.-morpholinoethyl)piperazinol)isoquinoline, m. 145-6.degree. (dil. EtOH). A mixt. of 12 g. I, 90 ml. Ac2O and 90 ml. HC(OEt)3 was refluxed 7 hrs. and the soln. cooled to yield 13 g. .alpha.-ethoxymethylenehomophthalimide (VII), m. 236-9.degree. (dil. MeOH). Hydrogenation of 5 g. VII in 200 ml. EtOH in the presence of 0.2 g. platinum oxide at atm. pressure yielded 3.4 g. .alpha.-methylhomophthalimide, m. 140-2.degree. (dil. EtOH). H2O2 (1 ml., 30%) was added to a soln. of 1.5 g. III in 6 ml. HOAc, after the exothermic reaction had subsided, 1 ml. concd. HCl added dropwise, the mixt. kept 1 hr. and treated with ice water to yield 1.5 g. 2-methyl-3,4-dichlorohomophthalimide (VIII), m. 137-8.degree. (EtOH). The structure of VIII was confirmed by N.M.R. spectra. A mixt. of 3 g.
3-chloro-N-methyl-homophthalimide (IX), 30 ml. dioxane and 8.4 ml. concd. HCl was heated at 85.degree. (oil-bath), treated dropwise with 9 ml. H2O2, and cooled to yield 3.7 g. .alpha.,.alpha.-dichloro-N-methyl-homophthalimide, m. 149-51.degree. (dil. EtoH). Use of 1 g. 3,4-dichloro-N-methylisocarbostyril in place of IX as above yielded 1 g. .alpha.,.alpha.-dichloro-N-methylhomophthalimide, m. 149-50.degree.. Similarly, chlorination of 10 g. I yielded 13.6 g. .alpha.,.alpha.dichlorohomophthalimide (X), m. 164-8.degree. (dil. EtOH). The reaction

PATEL 09/852,850

of X with substituted anilines (1 hr. reflux) yielded the corresponding phthalonimide anils (XI). The following XI were prepd. (R, Rl, m.p., and % yield given): H, 4-diethylaminophenyl, 195-6.degree. (C6H6-hexane), 64.3; H, 4-methoxyphenyl, 204-7.degree. (C6H6-hexane), 89.3; H, 4-chlorophenyl, 267-9.degree. (C6H6-hexane), 91.2; Me, NH2, 160-2.degree. (HOAc-H2O), 94.0; Me, NHPh, 176-7.degree. (EtOH-H2O), 71.5. X and XI reacted with o-phenylenediamine (45 hrs. reflux in C6H6) to yield, resp., quinoxalinoisocarbostyrils (XIIa), m. 265.degree. (HOAc) and XIIb, 203-5.degree. (HOAc). Secondary bases like morpholine reacted with X to give iminium salts (XIII), which were very hygroscopic and on catalytic hydrogenation led to hydrogenolytic cleavage to yield homophthalimides, while redn. with NaBH4 gave rise to water-sol. compds., from which no definite product could be isolated.

18630-91-6P

RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of)

18630-91-6 HCAPLUS

Isoquinoline, 1-morpholino-3-[4-(2-morpholinoethyl)-1-piperazinyl]- (8CI) (CA INDEX NAME)

L30 ANSWER 33 OF 35 HCAPLUS COPYRIGHT 2001 ACS

AN 1968:410341 HCAPLUS

69:10341 DN

ТI Synthesis of biologically interesting isoquinolines

ΑU Nair, Mohann D.

CS Ciba Res. Centre, Bombay, India

Symp. Syn. Heterocycl. Compounds Physiol. Interest, Hyderabad, India, 1964 (1966), Meeting Date 1964, 107-13 CODEN: 16VOA6

Conference

English LA

For diagram(s), see printed CA Issue. GΙ

The Gabriel rearrangement of 4,4-dimethylhomophthalimide with POC13 gave as the major product 1-chloro-3-chloromethyl-4-methylisoquinoline (I), and as byproducts, 1-chloro-3-methyl-4-chloromethylisoquinoline, 1-chloro-3,4-dimethylisoquinoline, .alpha.-chloromethylhomophthalimide, and N-(3,4-dimethyl-1-isoquinolyl)-4,4-dimethylhomophthalimide. Nitration of I gave a 5-nitro deriv., which readily reacted with primary and secondary amines. An optimum yield of 62% in the rearrangement was obtained by adding a small amt. of water to the reaction mixt. prior to heating to 200.degree.. Rearrangement of the corresponding 4,4-diethyland 4,4-dipropylhomophthalimides gave 1-chloro-3-(.beta.-chloroethyl)-4ethylisoquinoline and 1-chloro-3-(2-chloropropyl)-4-propylisoquinoline, resp. 4-Alkyl-4-benzylhomophthalimides were prepd. by hydrogenating 4-benzylidenehomophthalimide over PtO2, and then treating with an alkyl iodide. The 4-Me, 4-Et, and 4-Pr derivs. obtained were treated with POC13, giving C-debenzylation in all cases. The 4-Me compd. gave 1,3-dichloro-4-methylisoquinoline, while the 4-Et and 4-Pr derivs. gave isocarbostyril derivs. Some of the compds. showed borderline biol. activities. The most active was 4-methyl-1-morpholino-3-(morpholinomethyl)-isoquinoline, which showed high antitussive activity and was well tolerated.

15896-93-2P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 15896-93-2 HCAPLUS

1,3(2H,4H)-Isoquinolinedione, 2-(3,4-dimethyl-1-isoquinolyl)-4,4-dimethyl-

_ (8CI) - (CA INDEX NAME)

L30 ANSWER 34 OF 35 HCAPLUS COPYRIGHT 2001 ACS

1967:482075 HCAPLUS

67:82075

Isoquinolines. I. Rearrangement of .alpha.,.alpha.-dialkylhomophthalimides to 1-chloro-3, 4-dialkylisoquinoline derivatives

ΑU Marquardt, Fritz H.; Nair, Mohann D.

CIBA, Goregaon, India CS

Helv. Chim. Acta (1967), 50(6), 1469-76 CODEN: HCACAV SO

DΨ Journal

LA German

GI

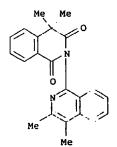
For diagram(s), see printed CA Issue. On reinvestigation of the reaction of wet POC13 with .alpha.,.alpha.dimethylhomophthalimide, 1-chloro-3-chloromethyl-4-methylisoquinoline (I) and 1-chloromethyl-3-methylisoquinoline were isolated as the main products (aside from some substances resulting from a redox disproportionation). The production of these two substances can be rationalized by assuming a mechanism in which the rarrangement product is a protonated deriv. of 3,4-dimethylene-3,4-dihydroisoquinoline. With .alpha.,.alpha.diethylhomophthalimide, the only isolated product was a deriv. of 1-chloro-3,4-diethylisoquinoline, with a Cl atom in .beta.-position to one of the Et groups, while with .alpha.-methyl-.alpha.-benzylhomophthalimide, the isolated product was 1,3-dichloro-4-methylisoquinoline, i.e. elimination occurred instead of rearrangement. Also these results are in agreement with the proposed mechanism.

ΙT 15896-93-2P

RL: SPN (Synthetic preparation); PREP (Preparation)

(prepn. of) 15896-93-2 HCAPLUS RN

1,3(2H,4H)-Isoquinolinedione, 2-(3,4-dimethyl-1-isoquinolyl)-4,4-dimethyl-(8CI) (CA INDEX NAME)



L30 ANSWER 35 OF 35 HCAPLUS COPYRIGHT 2001 ACS 1967:421848 HCAPLUS

DN -

67:21848

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New antitussive isoquinoline derivatives
TI
       CIBA Ltd.
PA
      Fr. M., 10 pp.
CODEN: FMXXAJ
SO
DT
       Patent
       French
FAN.CNT 1
                                                              APPLICATION NO.
       PATENT NO.
                               KIND DATE
                                                                                      DATE
                                        19660131
       FR 3782
                                        19630121
PRAI CH
                                        19640121
       CH
       For diagram(s), see printed CA Issue.
GΙ
       New antitussive isoquinoline derivs. with general formula (I) are prepd.
AB
       A mixt. of 9 g. 1-chloro-3-chloromethyl-4-methylisoquinoline (III) and 40 cc. piperidine (III) is heated in a sealed tube 8 hrs. at 150.degree., the
       reaction mixt. concd. in vacuo, treated with water, and extd. with CH2C12,
       the ext. dried and evapd. to dryness, and the residue in CHCl3 passed
       through activated alumina to give 4-methyl-1-piperidino-3-
       piperidinomethylisoquinoline, m. 111.degree. (water-EtOH).
       products are prepd. in a similar way (starting materials, reaction time, reaction temp., final product, m.p., derivs., and m.p. given): II (9 g.), pyrrolidine (40 cc.), 8 hrs., 150.degree., 4-methyl-1-(1-pyrrolidinyl)-3-
       (1-pyrrolidinylmethyl)isoquinoline, -, hydrochloride, 239.degree.; II (8
       g.), N-methylpiperazine (IV) (50 cc.), 8 hrs., 150.degree., 4-methyl-1-(N'-methylpiperazino)-3-(N'-methylpiperazinomethyl)isoquinoline
      4-methyl-1-(N'-methylpiperazino)-3-(N'-methylpiperazinomethyl)3, 110-11.degree., hydrochloride, 238.degree.; II (8 g.), N-(.beta.-hydroxyethyl)piperazine (40 cc.), 8 hrs., 150.degree., 4-methyl-1-[N'-(.beta.-hydroxyethyl)piperazino]-3-[N'-(.beta.-
       hydroxyethyl)piperazinomethyl)isoquinoline, 112.degree., hydrochloride, 262.degree. (decompn.); II (6 g.), Et2NH (15 cc.), 8 hrs., 150.degree.,
       4-methyl-1-diethylamino-3-diethylaminomethylisoquinoline, -, dimaleate,
       109-11.degree.; II (4.5 g.), ethanolamine (15 cc.), 3 hrs., 130.degree.
       4-methyl-1-(.beta.-hydroxyethylamino)-3-(.beta.
       hydroxyethylaminomethyl)isoquinoline, -, hydrochloride, 252-4.degree.; II
       (5 g.), N-carbethoxypiperazine (V) (20 cc.), 6 hrs., 140.degree.,
       4-methyl-1-(N'-carbethoxypiperazino)-3-(N'-carbethoxypiperazinomethyl)isoq
       uinoline, 90-2.degree., -, -; II (5 g.), 2-methylpiperidine (20 cc.), 6 hrs., 140.degree., 1-chloro-4-methyl-3-(2-methylpiperidinomethyl)isoquinol
       ine (VI), 106-8.degree., -, -; VI (6 g.), morpholine (VII) (20 cc.), 14 hrs., 170.degree., 4-methyl-1-morpholino-3-(2-
       methylpiperidinomethyl)isoquinoline, 103-4.degree., -,
       1-chloro-3-chloromethyl-4-methyl-5-nitroisoquinoline (VIII) (2 g.), VII
       (10 cc.), 2 hrs., 120.degree., 4-methyl-1-morpholino-3-morpholinomethyl-5-nitroisoquinoline (IX), 145-6.degree., -, -; VIII (2.5 g.), III (10 cc.), 2.5 hrs., 80.degree., 4-methyl-5-nitro-1-piperidino-3-
       piperidinomethylisoquinoline, 104-6.degree., -, -; VIII (2.5 g.), p-anisidine (4.55 g.), EtOH (80 cc.), 4 hrs., reflux, 1-p-anisidino-3-p-
       anisidinomethyl-4-methyl-5-nitroisoquinoline, 183-5.degree., -, -;
       1,7-dichloro-3-chloromethyl-4-methylisoquinoline (X) (4 g.), VII (50 cc.),
       4 hrs., reflux, 7-chloro-4-methyl-1-morpholino-3-
      morpholinomethylisoquinoline, 120.degree., maleate, -; VIII (5 g.), III (8 cc.), EtOH (75 cc.), 1 hr., reflux, 1-chloro-4-methyl-5-nitro-3-piperidinomethylisoquinoline, 67-79.degree., -, -; II (4.5 g.), III (15
       cc.), 2 hrs., 80.degree., 1-chloro-4-methyl-3-
       piperidinomethylisoquinoline, 79-80.degree., -, -; VIII (3.5 g.), IV (2.58 g.), EtOH (100 cc.), 2 hrs., reflux, 1-chloro-3-(N'-
       methylpiperazinomethyl)-4-methyl-5-nitroisoquinoline, 173-5.degree., -, -;
       VIII (4 g.), V (10 cc.), EtOH (75 cc.), 1 hr., reflux,
       1-chloro-3-(N'-carbethoxypiperazinomethyl)-4-methyl-5-nitroisoquinoline,
       127-8.degree., -, -; VIII (2.71 g.), diethanolamine (4.5 g.), dioxane (50 cc.), 3 hrs., reflux, 1-chloro-3-[bis(.beta.-hydroxyethyl)aminomethyl)-4-
       methyl-5-nitroisoquinoline, 110-12.degree., -, -; II (5.0 g.), 4-methylpiperidine (5.5 cc.), 2 hrs., 80.degree., 1-chloro-3-(4-
       methylpiperidinomethyl)-4-methylisoquinoline, 83-5.degree., -, -; II (5.0 g.), concd. aq. NH3 (80 cc.), hydrated CuSO4 (1.0 g.), 30 hrs.,
       140.degree., bis(1-chloro-4-methyl-3-isoquinolylmethyl)amine,
       131-2.degree., -, -; II (5.0 g.), N-(.gamma.-aminopropyl)morpholine (6.5
       g.), 2 hrs., 100.degree., N,N-bis(1-chloro-4-methyl-3-isoquinolylmethyl)-N-
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(.gamma.-morpholinopropyl)amine, 110-12.degree., -, -. Some starting materials and other products are prepd. as follows: II (6 g.) is added slowly with stirring to a cooled mixt. of 15 cc. concd. H2SO4 and 15 cc. fuming HNO3 and the mixt. stirred 1.5 hrs. below 5.degree. and poured over a mixt. of ice and water to ppt. VIII, m 104-5.degree. (EtOH). A mixt. of 4 g. IX, 0.3 g. Pd-C and 150 cc. 95% EtOH is hydrogenated 1.5 hrs. to give $\hbox{5-amino-4-methyl-1-morpholino-3-morpholinomethylisoquinoline (XI), m.}\\$ 134-5.degree. (EtOH). A soln. of 1.6 g. NaNO2 in 5 cc. water is added slowly to a cooled soln. of 8 g. XI in 6 cc. concd. HCl and 6 cc. water. the resulting soln. poured into a cooled soln. of Cu2Cl2 (prepd. from 8 g. CusO4) and then is heated at 60.degree., and the ppt. suspended in 25 cc. water, alkalinized, and extd. with CHC13 to give 5-chloro-4-methyl-1morpholino-3-morpholinomethylisoquinoline, m. 104.degree... 4,4-Dimethylhomophthalimide (15 g.) is added slowly with stirring to a cooled (-10.degree.) mixt. of 30 cc. concd. H2SO4 and 30 cc. fuming HNO3 and the mixt. stirred 1 hr. below 20.degree. and poured over a mixt. of ice and water to ppt. 4,4-dimethyl-7-nitrohomophthalimide (XII), m. 209-11.degree. (EtOH). A mixt. of 23.4 g. XII, 0.5 g. Pd-C, and 200 cc. MeOH is hydrogenated at 50.degree./3.4 atm. .appxx.1.5 hrs. to give 4,4-dimethyl-7-aminohomophthalimide (XIII), m. 176-9.degree. (MeOH) Concd. H2SO4 (26 g.) is added slowly to a mixt. of 20 g. XIII and 90 cc. water, and cooled at 0.degree., 8.4 g. NaNO2 in 24 cc. water added slowly to it, and this mixt. is added slowly to a soln. of Cu2Cl2 (prepd. from 33.4 g. CuSO4), and the mixt. heated at 60.degree. 30 min., cooled, dild. with water, and extd. with CHCl3 to give 4,4-dimethyl-7chlorohomophthalimide (XIV), m. 200.degree. (EtOH). A mixt. of 10 g. XIV, 0.5 cc. water, and 40 cc. POCl3 is heated in a sealed tube at 200.degree. 5 hrs. to give X, m. 135.degree. (hexane-CHCl3). Some recipes for the prepn. of pharmacol. compns. are also given. 14576-16-0P 14576-17-1P 14577-67-4P 14601-03-7P 14601-04-8P 14601-07-1P 14657-46-6P 14657-48-8P 14657-49-9P 14657-50-2P 14657-51-3P 14657-52-4P 14825-52-6P RL: SPN (Synthetic preparation); PREP (Preparation) (prepn. of) 14576-16-0 HCAPLUS 1-Piperazineethanol, 4-[{1-(4-(2-hydroxyethyl)-1-piperazinyl}-4-methyl-3-

isoquinolinyl)methyl)- (9CI) (CA INDEX NAME)

RN 14576-17-1 HCAPLUS

CN 1-Piperazinecarboxylic acid, 4-[[1-[4-(ethoxycarbonyl)-1-piperazinyl]-4-methyl-3-isoquinolinyl]methyl]-, ethyl ester (9CI) (CA INDEX NAME)

RN 14577-67-4 HCAPLUS

CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-((4-methyl-1-piperazinyl)methyl]- (8CI, 9CI) (CA INDEX NAME)

RN 14601-03-7 HCAPLUS

CN Isoquinoline, 4-methyl-1-(1-pyrrolidinyl)-3-(1-pyrrolidinylmethyl)-, hydrochloride (8CI) (CA INDEX NAME)

$$Me$$
 CH_2
 N

●x HCl

RN 14601-04-8 HCAPLUS

CN Isoquinoline, 4-methyl-1-(4-methyl-1-piperazinyl)-3-[(4-methyl-1-piperazinyl)methyl]-, hydrochloride (8CI) (CA INDEX NAME)

●x HCl

RN 14601-07-1 HCAPLUS
CN Isoquinoline, 7-chloro-4-methyl-1-morpholino-3-(morpholinomethyl)-,
maleate (8CI) (CA INDEX NAME)

СМ

CRN 47438-17-5 CMF C19 H24 C1 N3 O2

CM 2 .

CRN 110-16-7 CMF C4 H4 O4 CDES 2:2

Double bond geometry as shown.

RN 14657-46-6 HCAPLUS
CN Isoquinoline, 4-methyl-1-(1-piperidinyl)-3-(1-piperidinylmethyl)- (9CI)
(CA INDEX NAME)

- RN 14657-48-8 HCAPLUS
- Isoquinoline, 4-methyl-3-[(2-methyl-1-piperidinyl)methyl]-1-(4-morpholinyl)- (9CI) (CA INDEX NAME) CN

- RN 14657-49-9 HCAPLUS
- Isoquinoline, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-5-nitro-(9CI) (CA INDEX NAME) CN

- 14657-50-2 HCAPLUS
 Isoquinoline, 4-methyl-5-nitro-1-(1-piperidinyl)-3-(1-piperidinylmethyl)-(9CI) (CA INDEX NAME) RN CN

- 14657~51-3 HCAPLUS
- CN 5-Isoquinolinamine, 4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)-(9CI) (CA INDEX NAME)

RN 14657-52-4 HCAPLUS
CN Isoquinoline, 5-chloro-4-methyl-1-(4-morpholinyl)-3-(4-morpholinylmethyl)(9CI) (CA INDEX NAME)

RN ,14825-52-6 HCAPLUS
CN '1-Piperazineethanol, 4,4'-{methylene(4-methyl-3,1-isoquinolinediyl)}di-,
hydrochloride (8CI) (CA INDEX NAME)

•x HCl